

FIG. 1

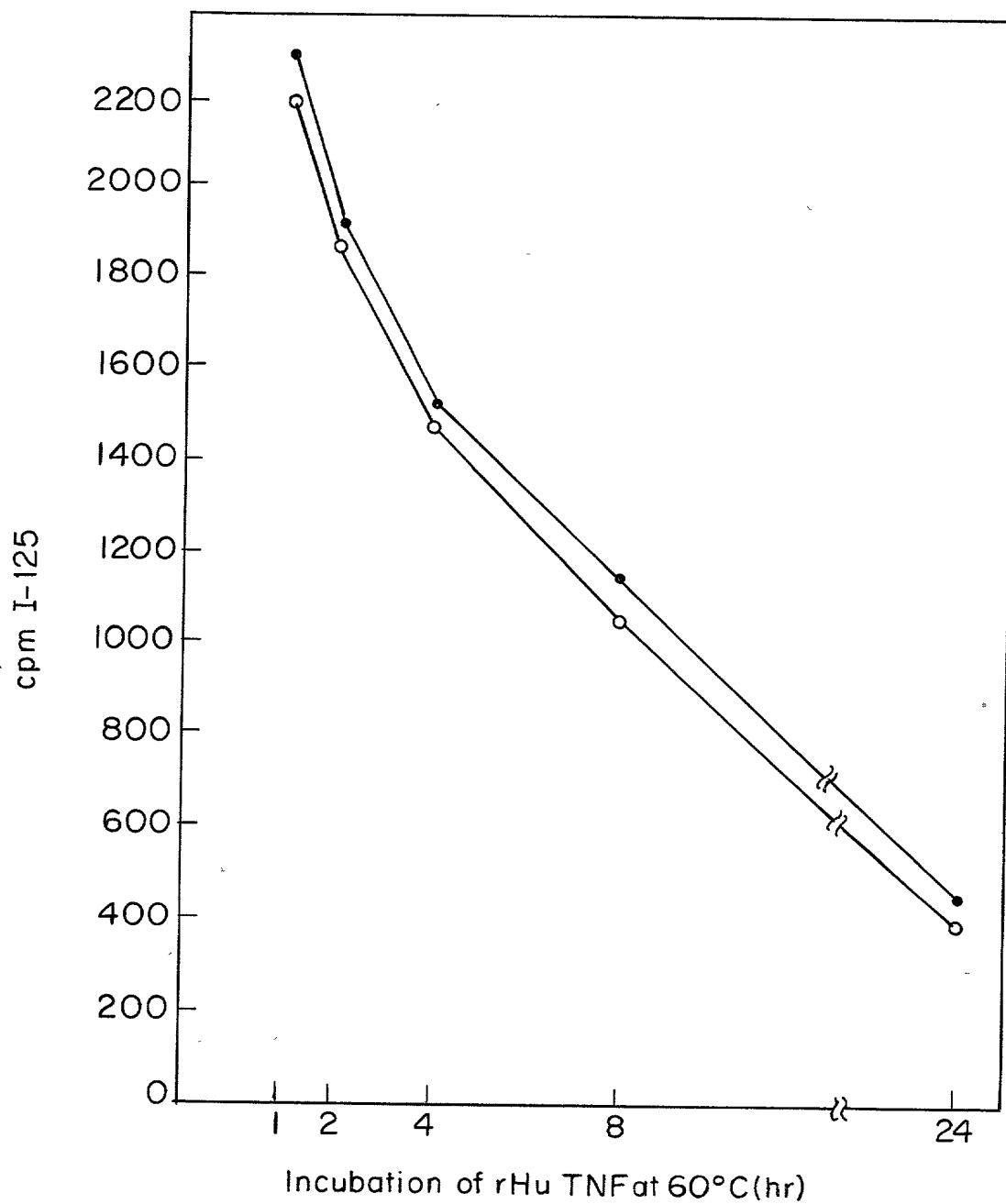


FIG. 2

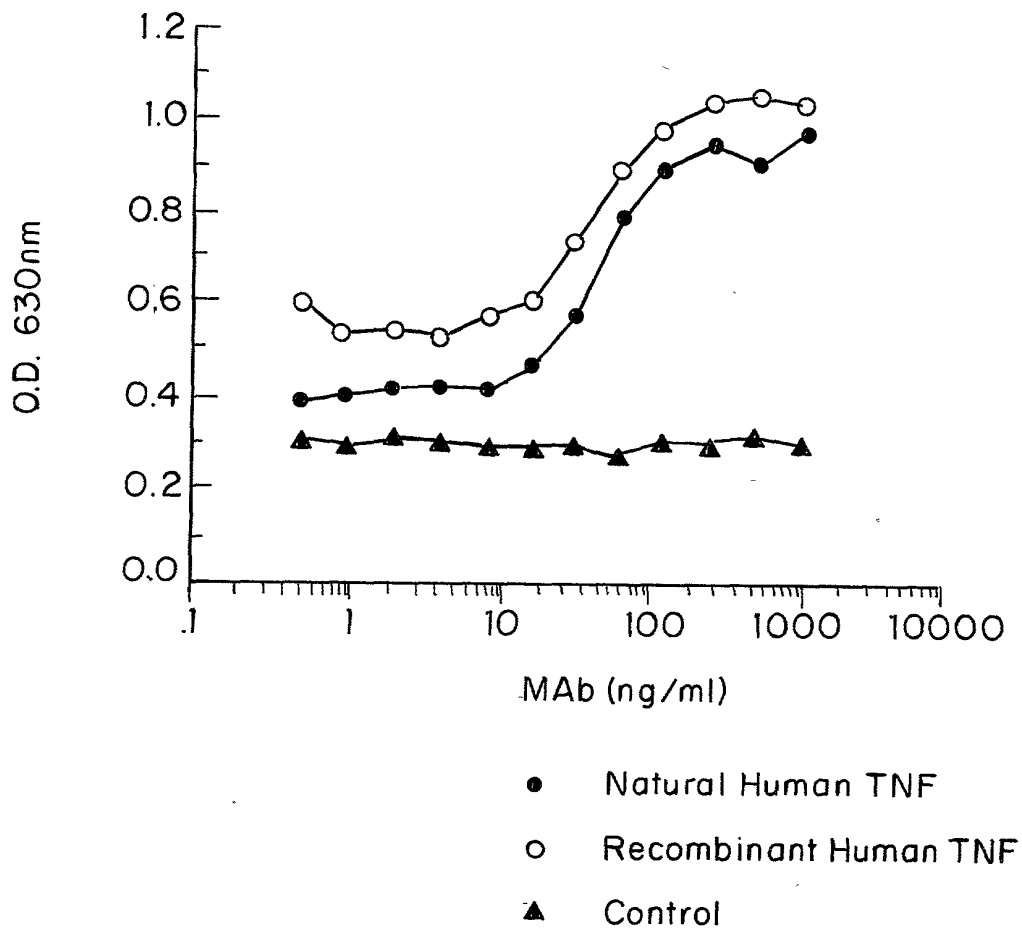


FIG. 3

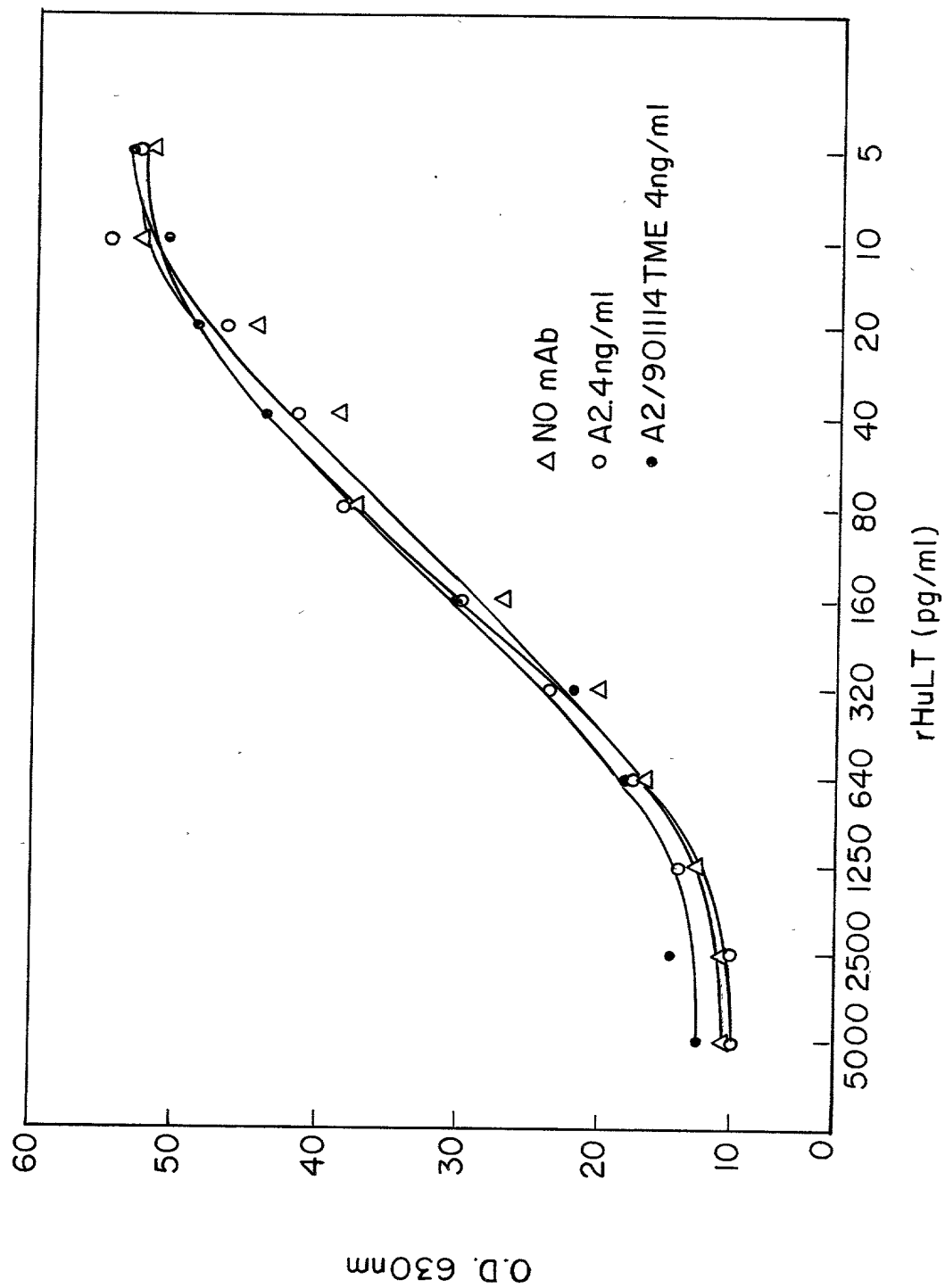


FIG. 4

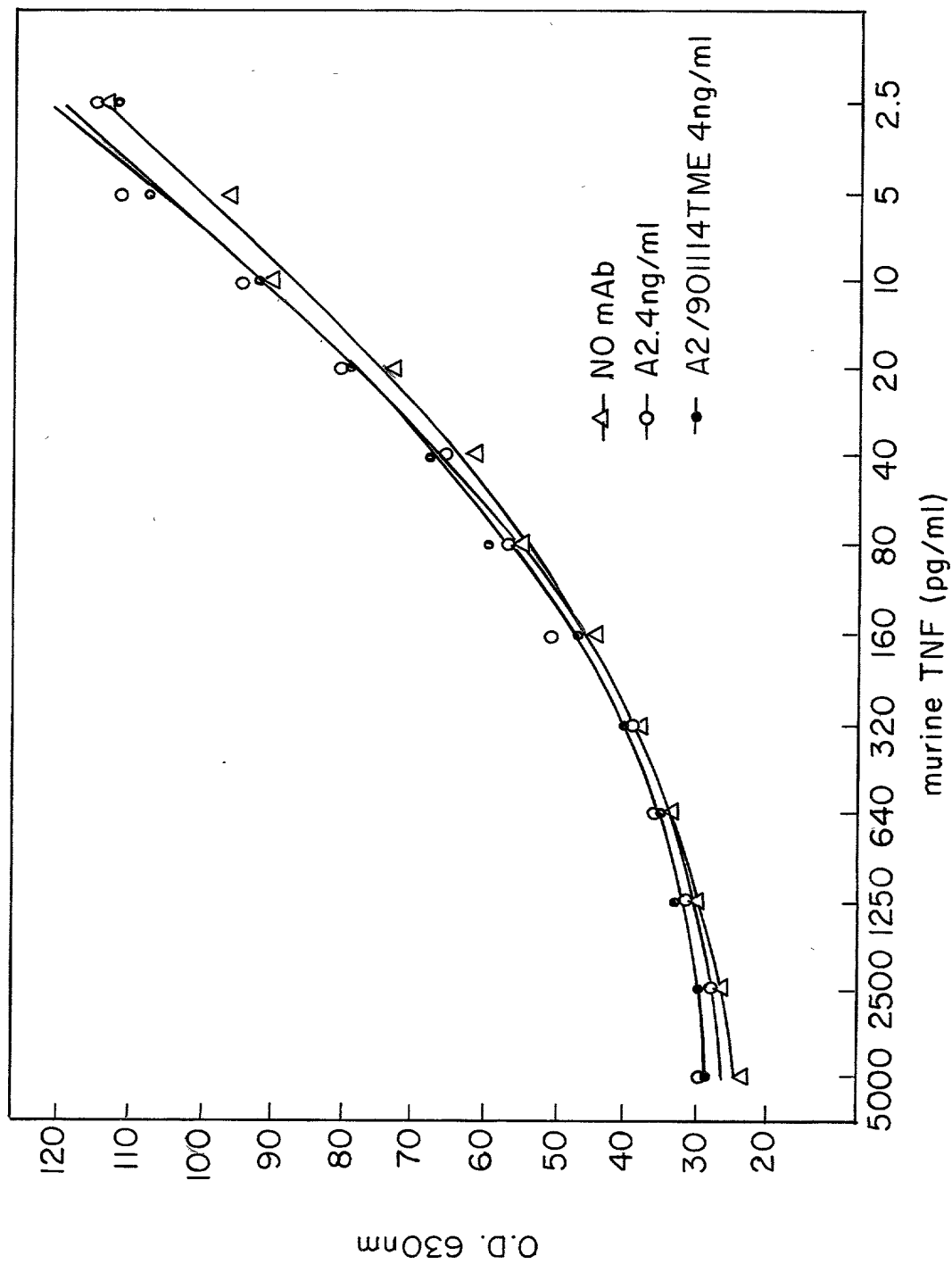


FIG. 5

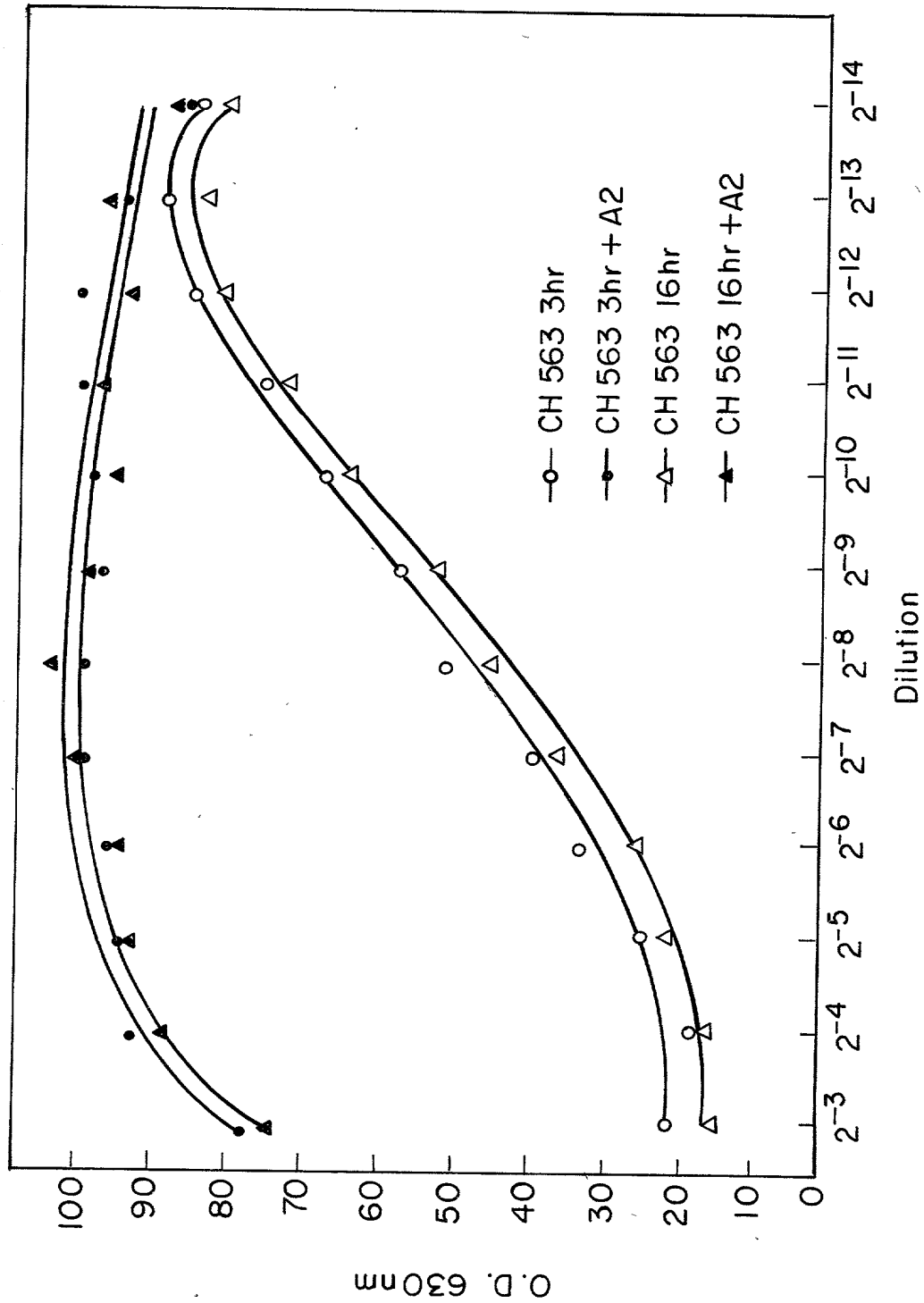


FIG. 6

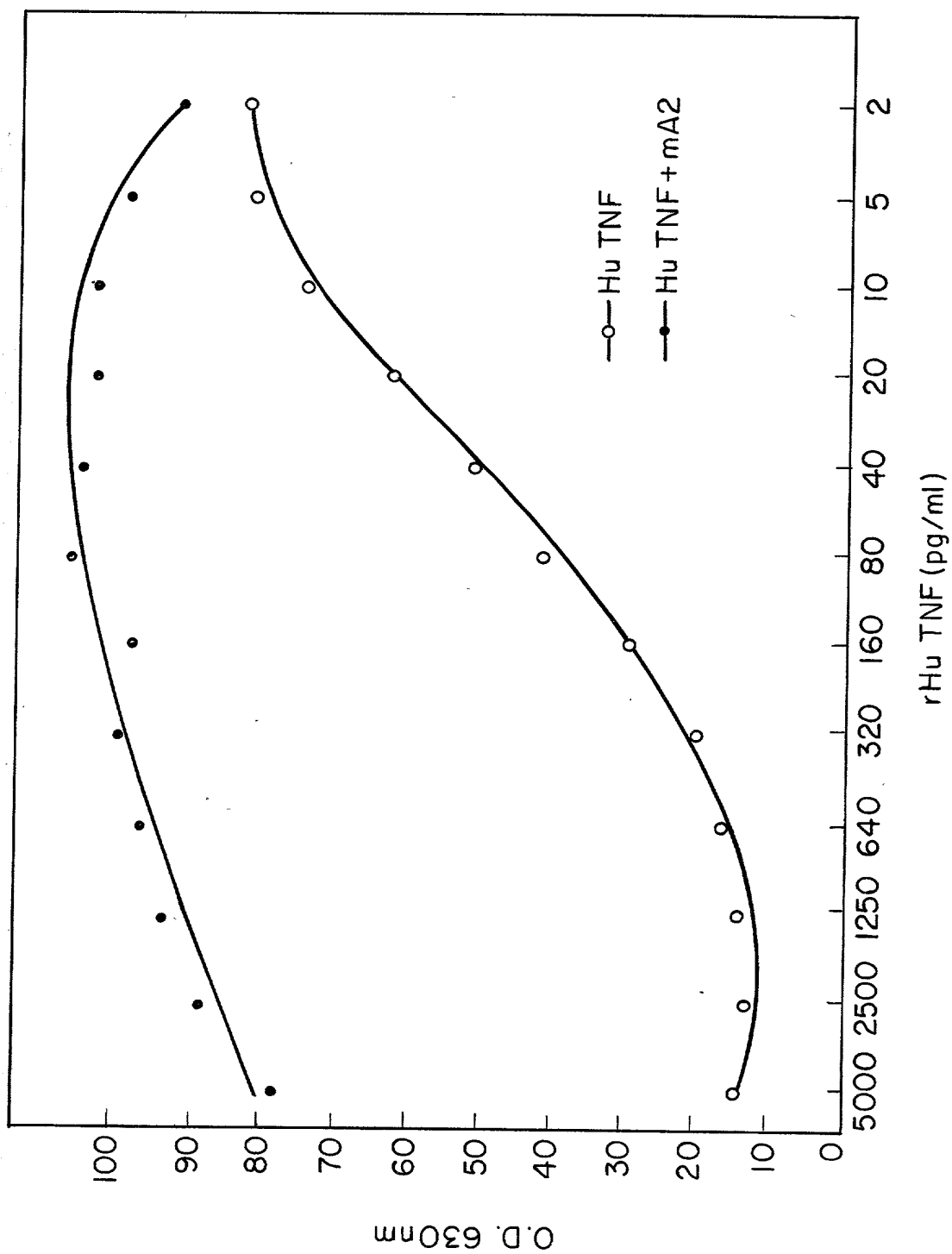


FIG. 7

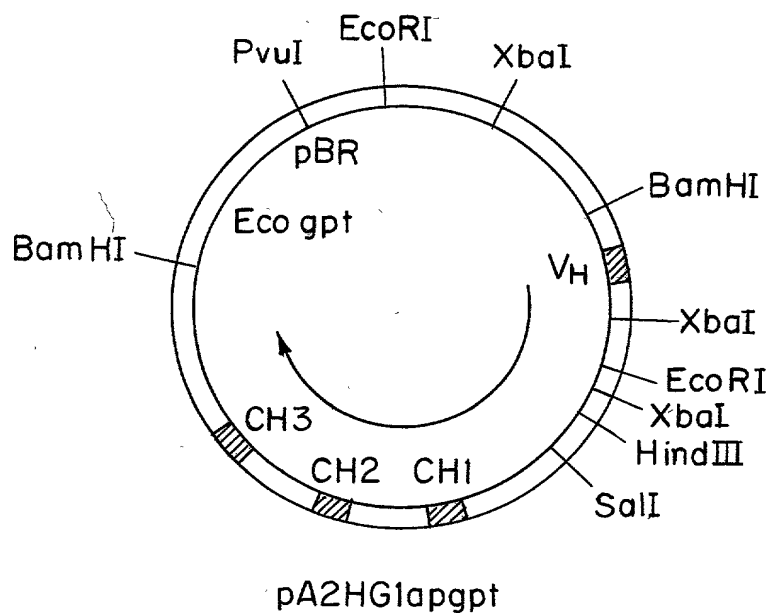


FIG. 8A

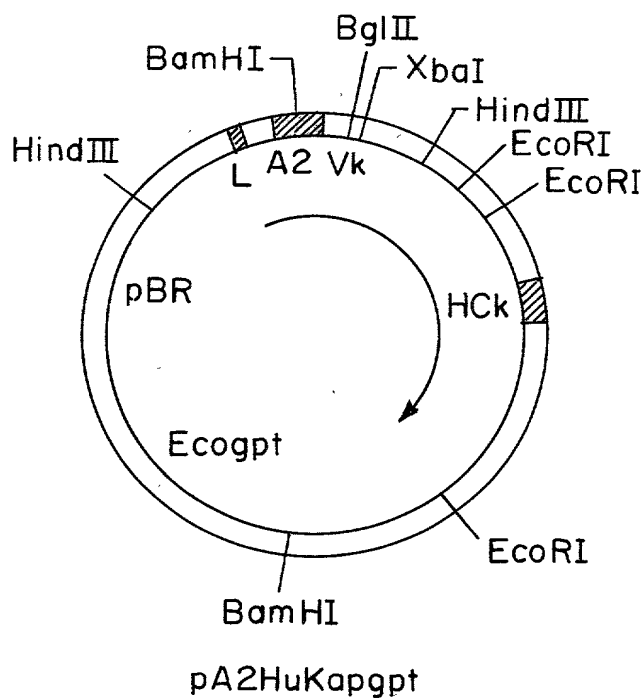


FIG. 8B

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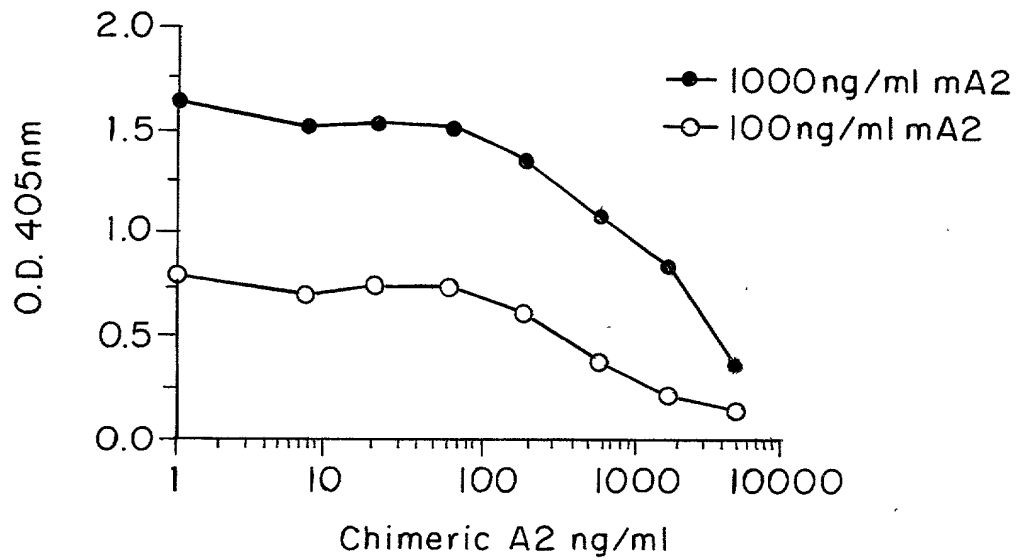


FIG. 9A

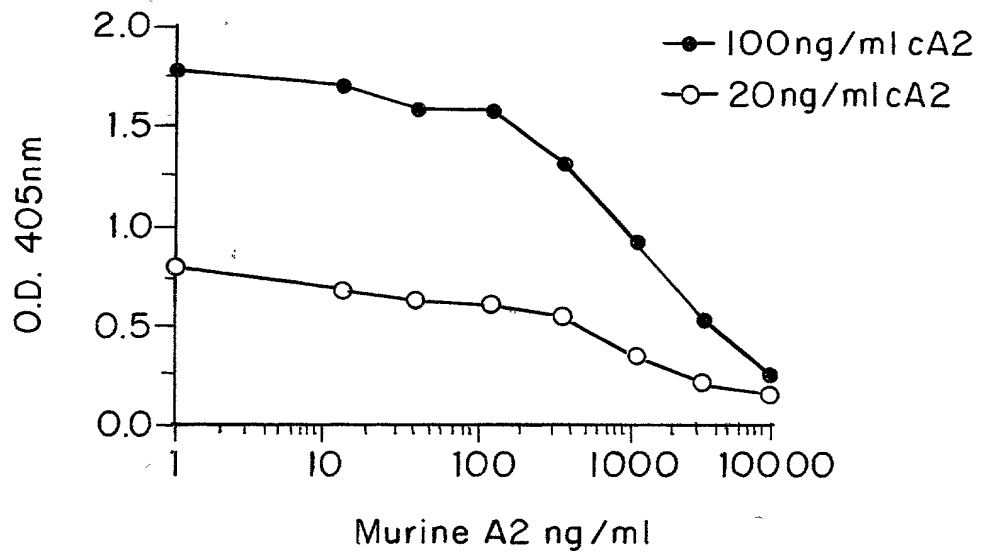


FIG. 9B

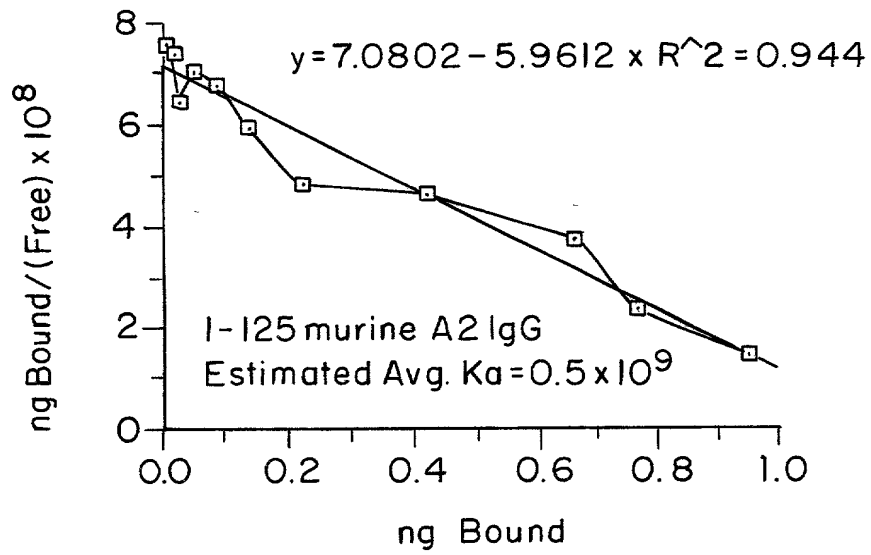


FIG. 10A

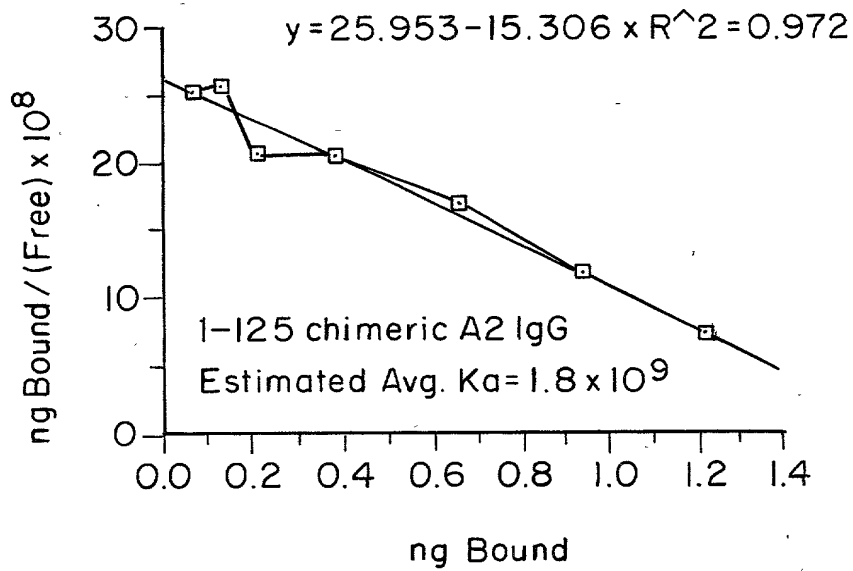


FIG. 10B

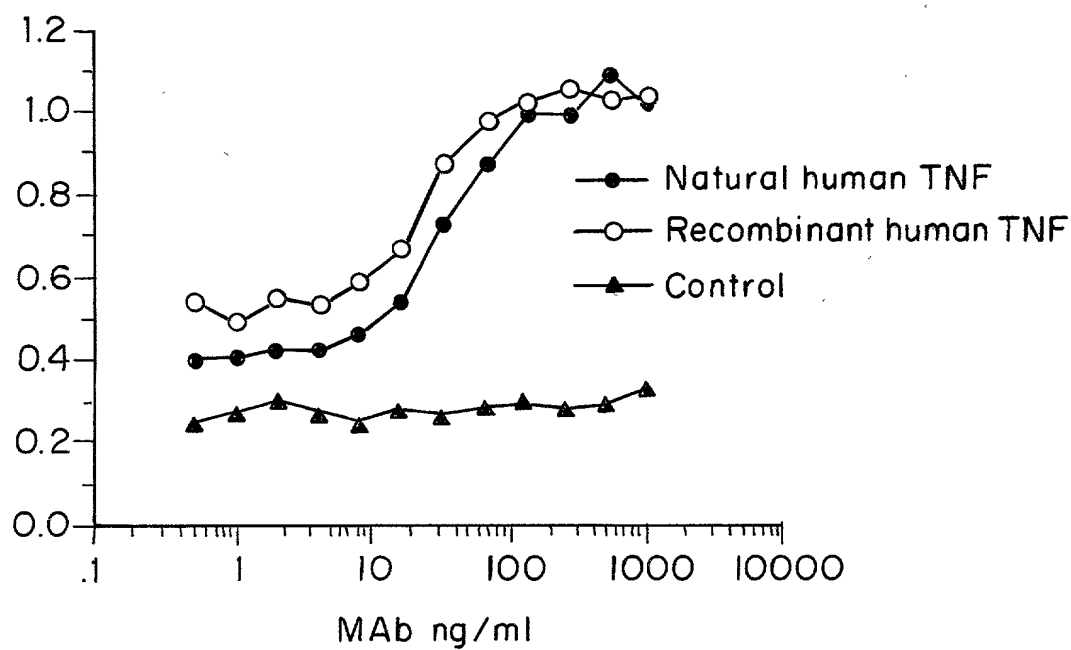


FIG. 11

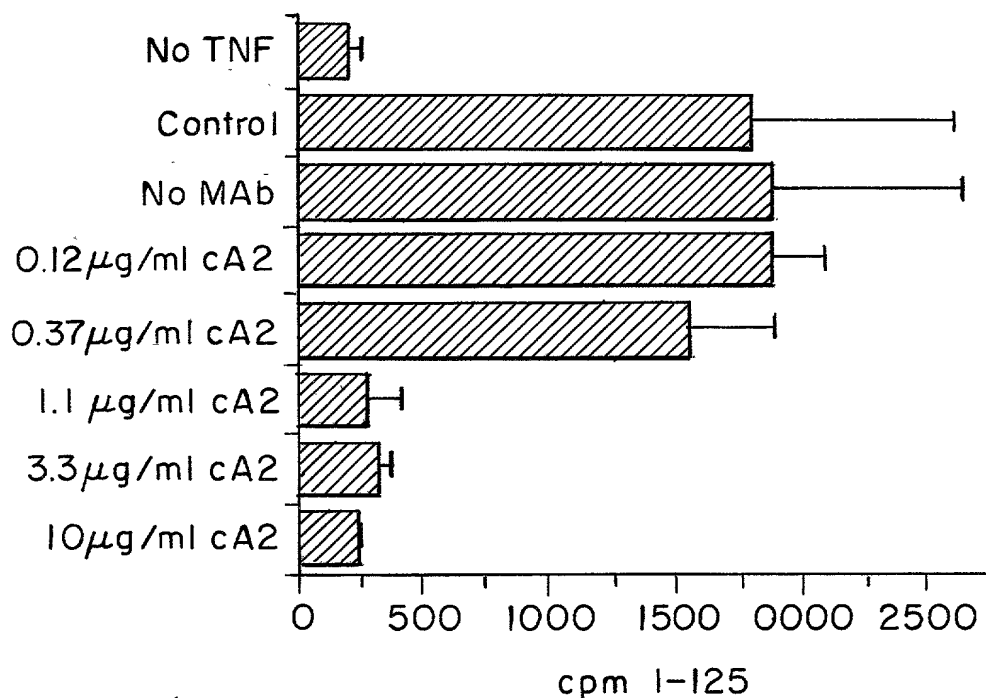


FIG. 12

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1 Val Arg Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His Val Val Ala Asn Pro
21 Gln Ala Glu Gly Gln Leu Gln Trp Leu Asn Arg Arg Ala Asn Ala Leu Leu Ala Asn Gly
41 Val Glu Leu Arg Asp Asn Gln Leu Val Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser
61 Gln Val Leu Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr Ile
81 Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser Ala Ile Lys Ser Pro
101 Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala Lys Pro Trp Tyr Glu Pro Ile Tyr Leu
121 Gly Gly Val Phe Gln Leu Glu Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp
141 Tyr Leu Asp Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu

FIG. 13

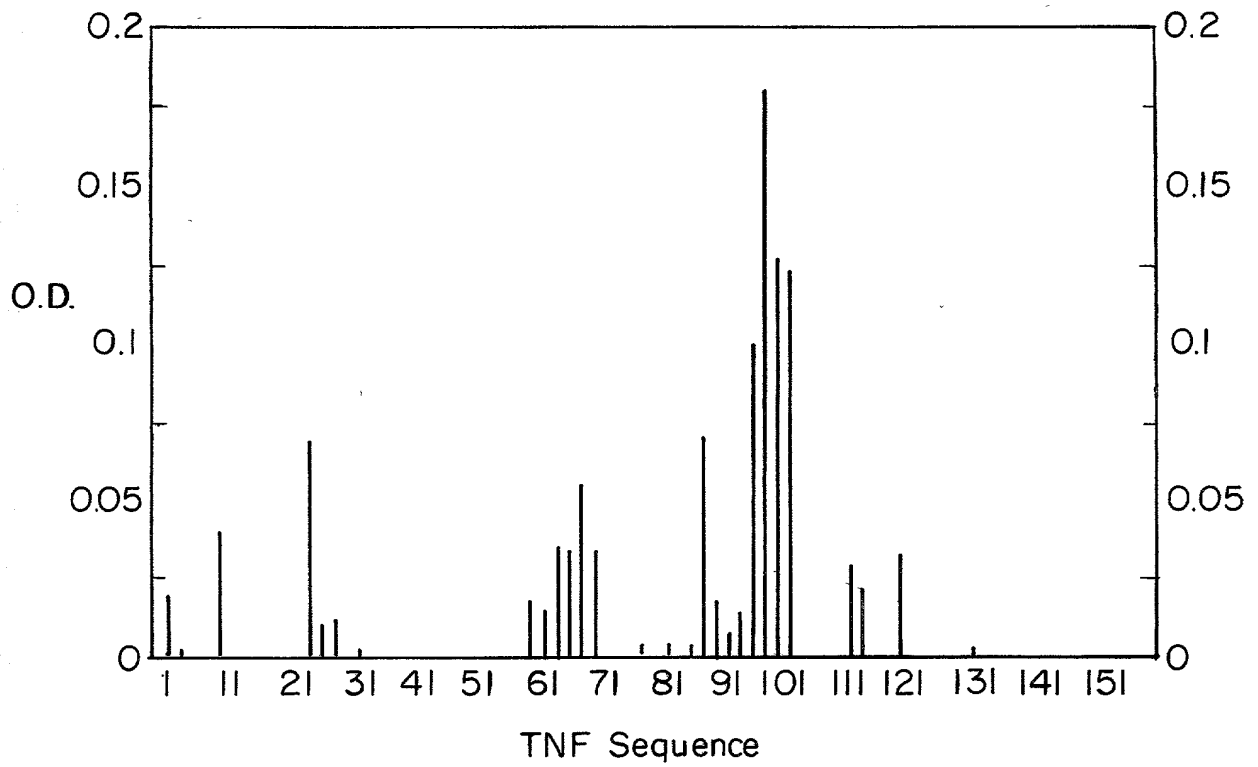


FIG. 14A

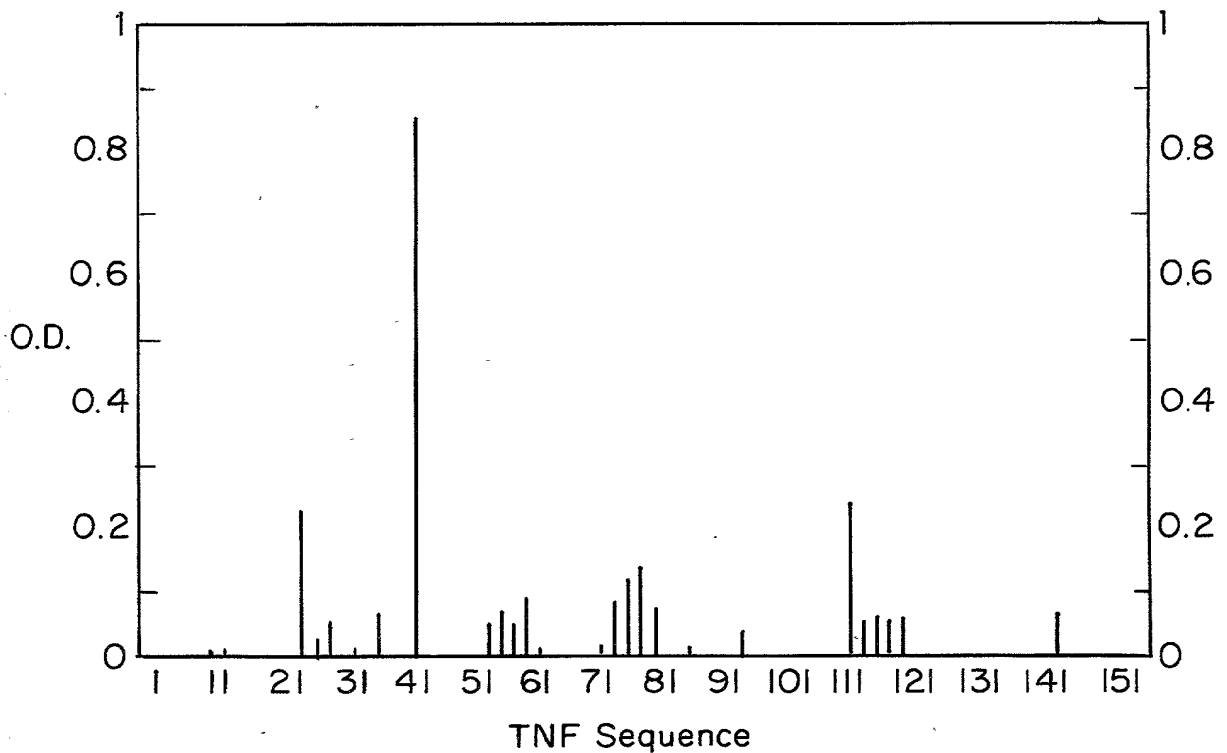


FIG. 14B

FIG. 15

1 Val Arg Ser Ser Arg Thr Pro Ser Asp Lys Pro Val Ala His Val Ala Asn Pro
10
21 Gln Ala Glu Gly Gln Leu Trp Leu Asn Arg Arg Ala Asn Ala Leu Leu Ala Asn Gly
30
41 Val Glu Leu Arg Asp Asn Gln Leu Val Pro Ser Glu Gly Leu Tyr Leu Ile Tyr Ser
50
61 Gln Val Leu Phe Lys Gly Gln Gly Cys Pro Ser Thr His Val Leu Leu Thr His Thr Ile
70
81 Ser Arg Ile Ala Val Ser Tyr Gln Thr Lys Val Asn Leu Leu Ser Ala Ile Lys Ser Pro
90
101 Cys Gln Arg Glu Thr Pro Glu Gly Ala Glu Ala Lys Pro Trp Tyr Glu Pro Ile Tyr Leu
110
121 Gly Gly Val Phe Gln Leu Glu Lys Gly Asp Arg Leu Ser Ala Glu Ile Asn Arg Pro Asp
130
141 Tyr Leu Asp Phe Ala Glu Ser Gly Gln Val Tyr Phe Gly Ile Ile Ala Leu
150

FIG. 15

GACATCTTGCTGACTCAGTCTCCAGCCATCCTGTCTGTGAGTCCAGGAGAAAGAGTCAGT
AspIleLeuLeuThrGlnSerProAlaIleLeuSerValSerProGlyGluArgValSer
TTCTCCTGCAGGGCCAGTCAGTTCGTTGGCTCAAGCATCCACTGGTATCAGCAAGAACA
PheSerCysArgAlaSerGlnPheValGlySerSerIleHisTrpTyrGlnGlnArgThr
AATGGTTCCTCAAGGCTTCTCATAAAGTATGCTTCTGAGTCTATGTCTGGATCCCTTCC
AsnGlySerProArgLeuLeuIleLysTyrAlaSerGluSerMetSerGlyIleProSer
AGGTTTAGTGGCAGTGGATCAGGGACAGATTTTACTCTTAGCATCAACACTGTGGAGTCT
ArgPheSerGlySerGlySerGlyThrAspPheThrLeuSerIleAsnThrValGluSer
GAAGATATTGCAGATTATTACTGTCAAGAAAGTCATAGCTGGCCATTACGTTTCGGCTCG
GluAspIleAlaAspTyrTyrCysGlnGlnSerHisSerTrpProPheThrPheGlySer
GGGACAAAATTGGAAAGTAAAA
GlyThrAsnLeuGluValLys

FIG. 16A

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GAAGTGAAGCTTGAGGAGTCTGGAGGAGGCTTGTTGTCACCTGGAGGATCCATGAACCTC
GluValLysLeuGluSerGlyGlyGlyLeuValGlnProGlyGlySerMetLysLeu
TCCTGTGTTGCCCTCTGGATTTCATTTTCAGTAACCACTGGATGAACCTGGGTCCGCCAGTCT
SerCysValAlaSerGlyPheIlePheSerAsnHisTrpMetAsnTrpValArgGlnSer
CCAGAGAAGGGCTTGAGTGGGTGCTGAAATTAGATCAAAATCTATTAAATTTCTGCAACA
ProGluLysGlyLeuGluTrpValAlaGluIleArgSerLysSerIleAsnSerAlaThr
CATTATGCCGAGTCTGTGAAAGGGAGGTTCAACCATCTCAAGAGATGATTCCAAAGTGCT
HisTyrAlaGluSerValLysGlyArgPheThrIleSerArgAspSerLysSerAla
GTGTACCTGCAAAATGACCGACTTAAGAACTGAAGACACTGGCGTTTATTACTGTCCAGG
ValTyrLeuGlnMetThrAspLeuArgThrGluAspThrGlyValTyrTyrCysSerArg
AATTACTACGGTAGTACCTACGACTACTGGGGCCCAAGCACCACTCTCACAGTGTCC
AsnTyrTyrGlySerThrTyrAspTyrTrpGlyGlnGlyThrThrLeuThrValSer

FIG. 16B

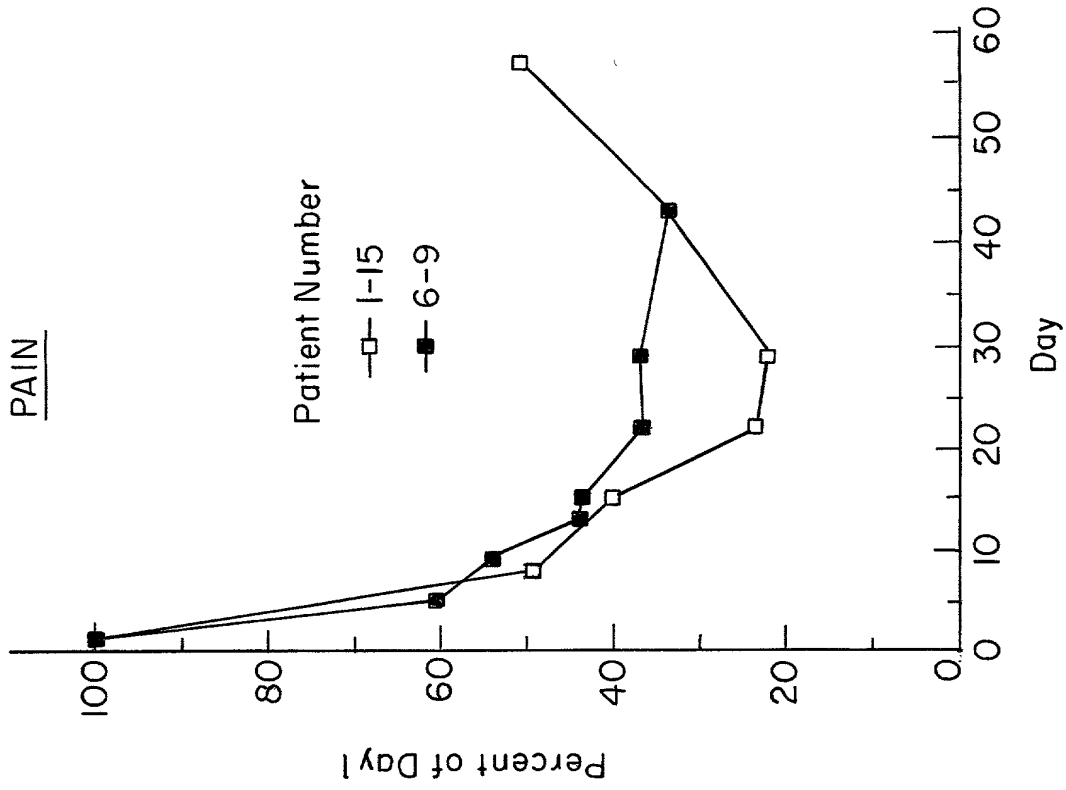


FIG. 18

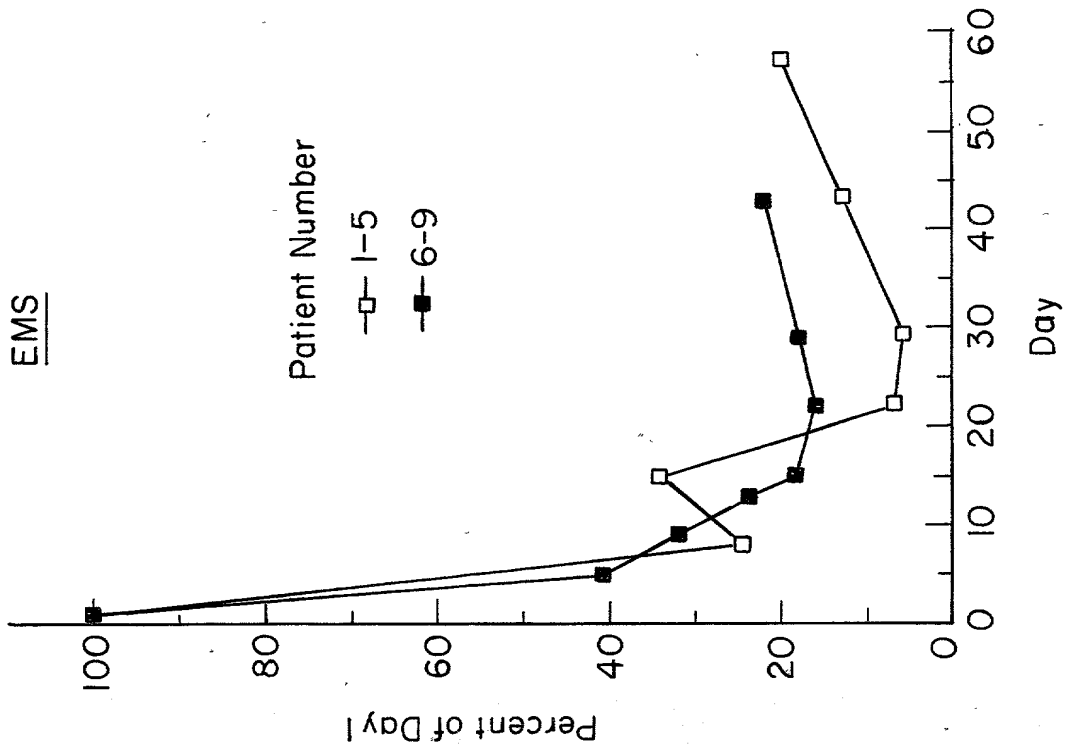


FIG. 17

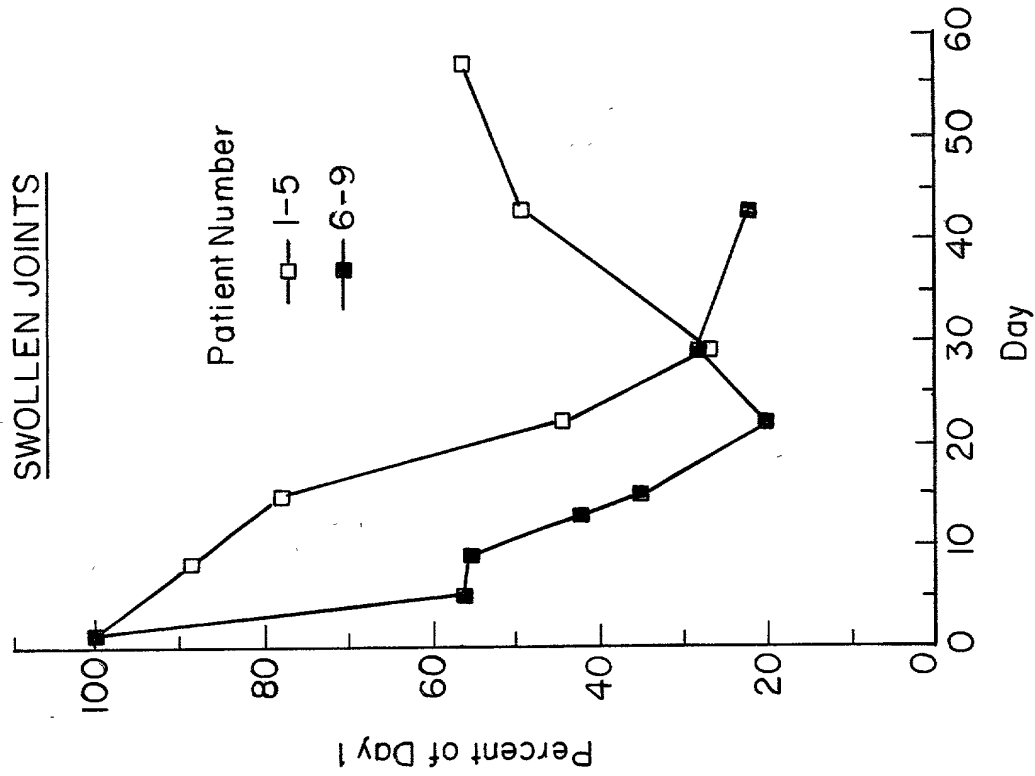


FIG. 20

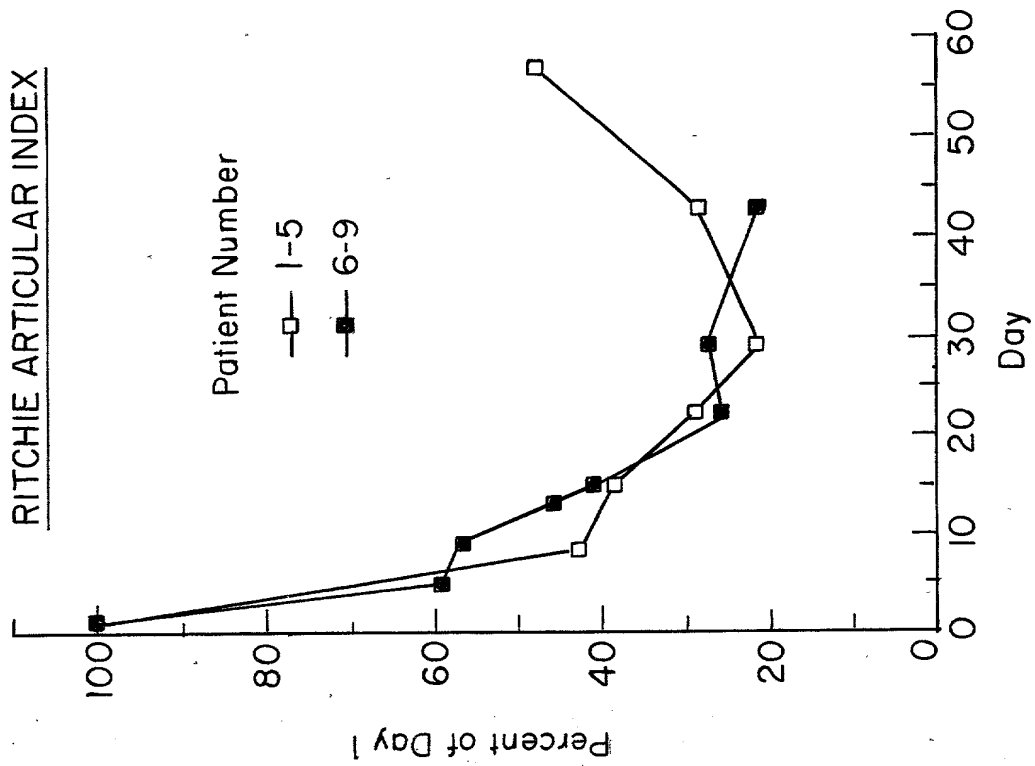


FIG. 19

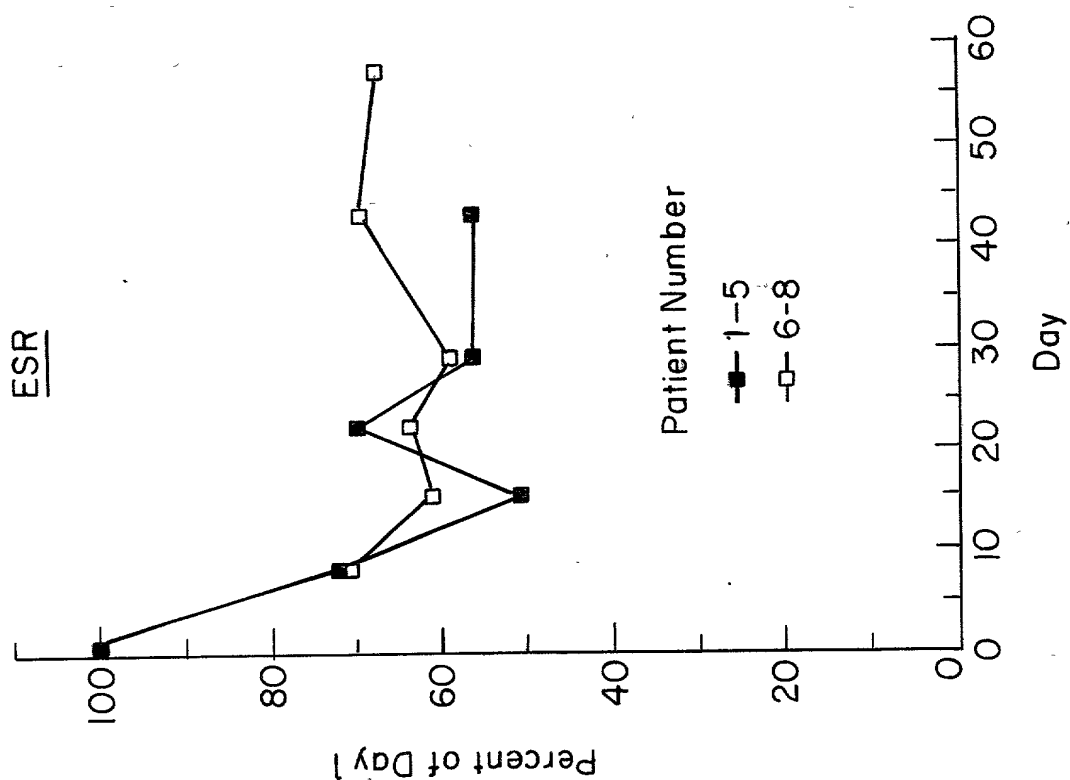


FIG. 22

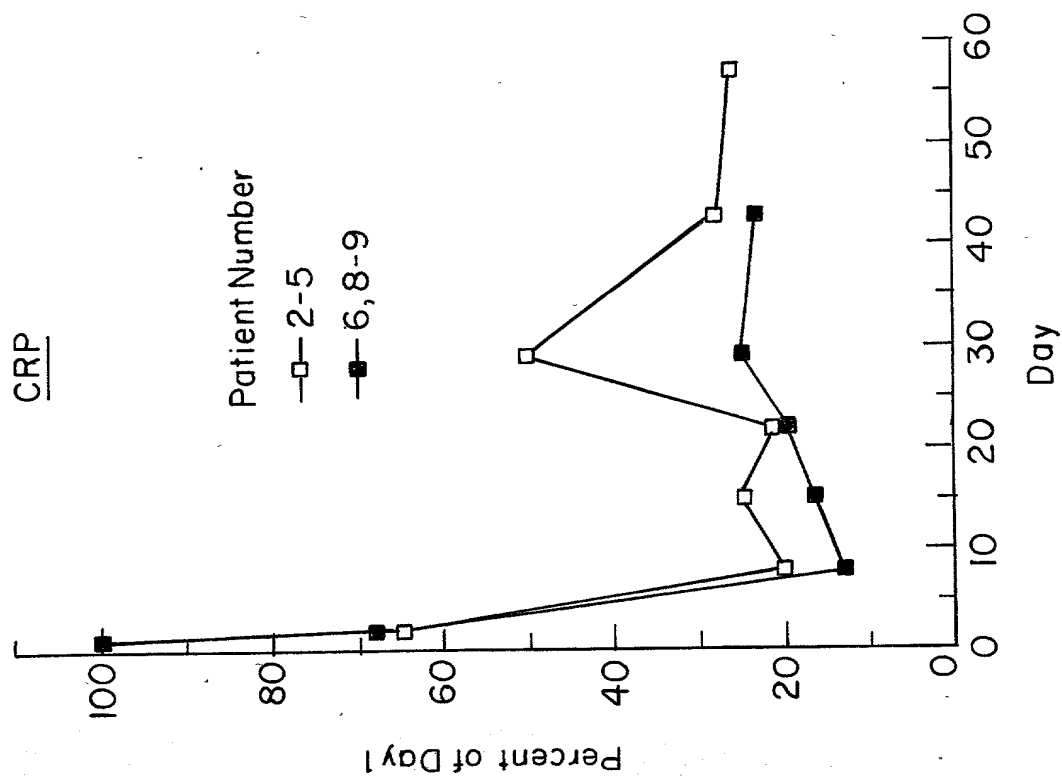


FIG. 21

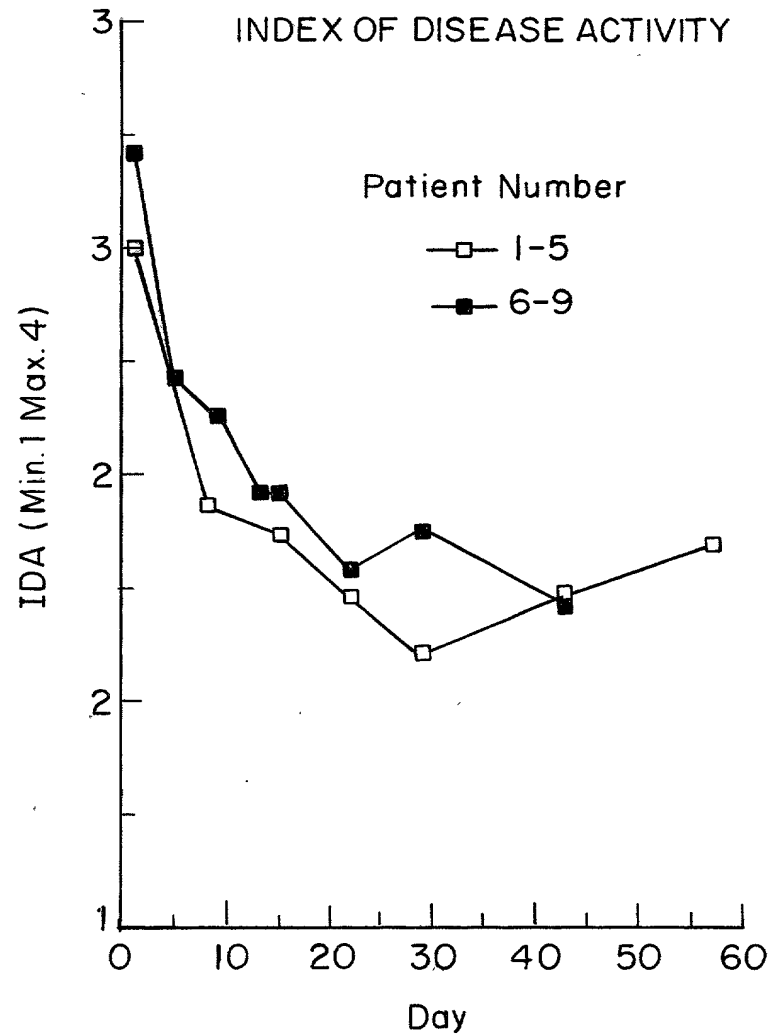
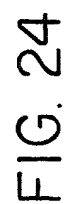


FIG. 23



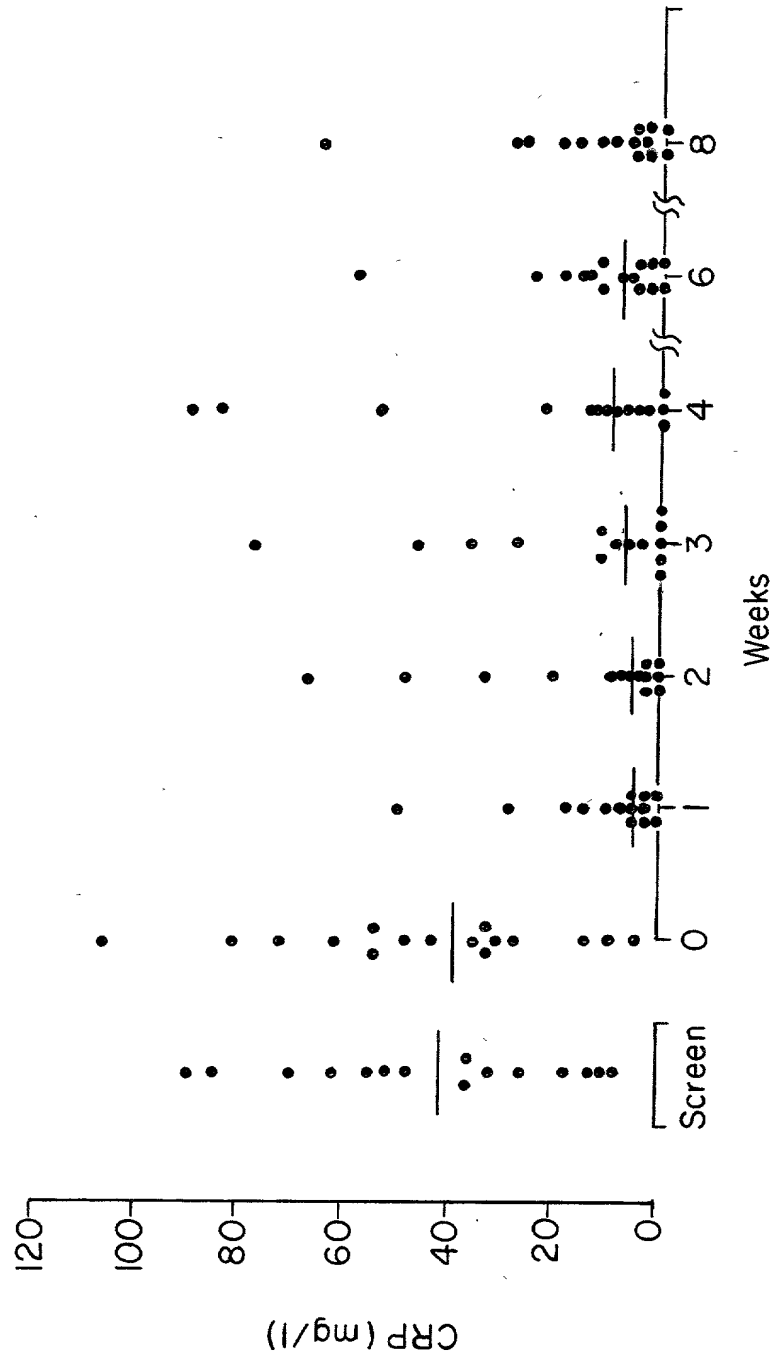
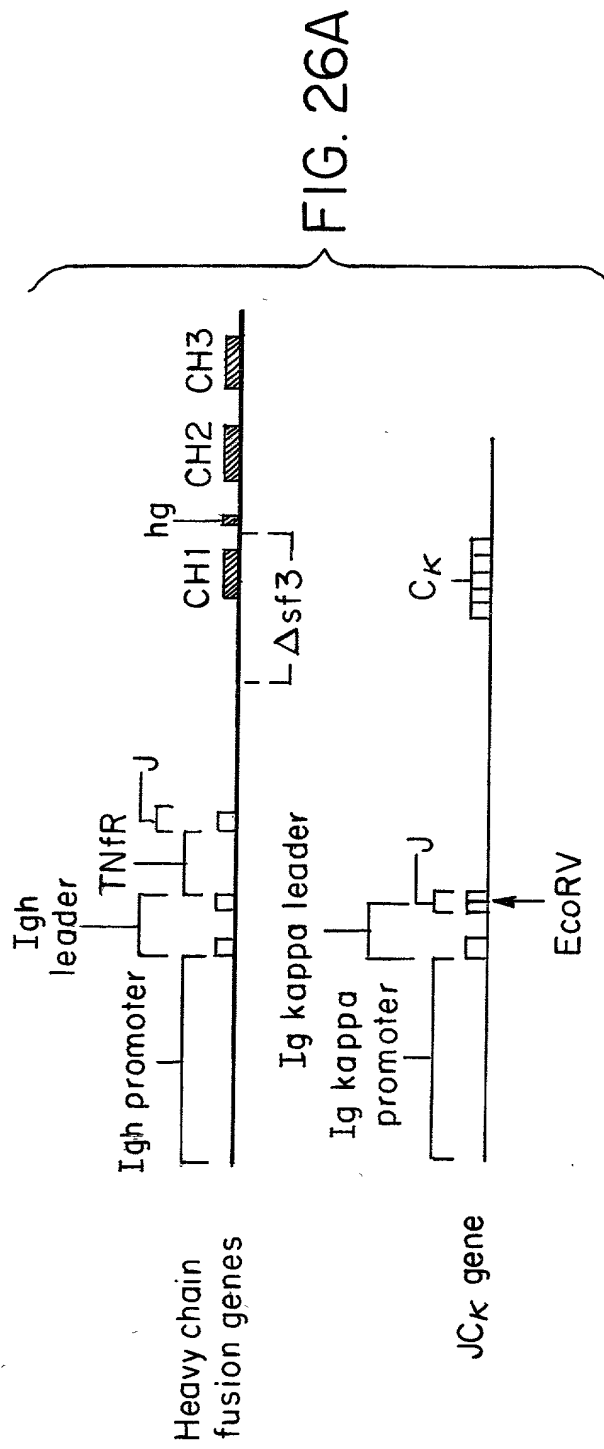
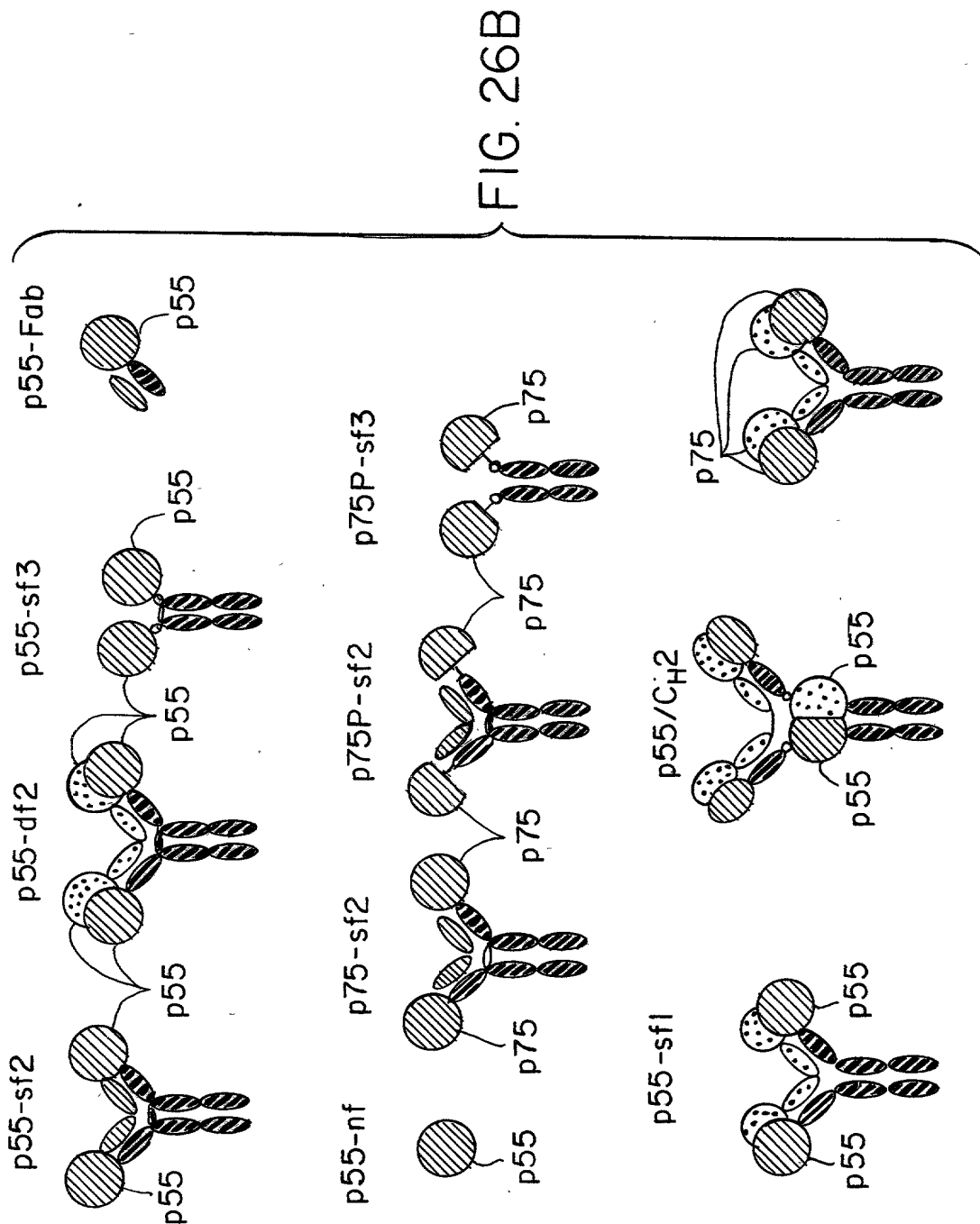
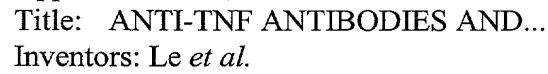


FIG. 25





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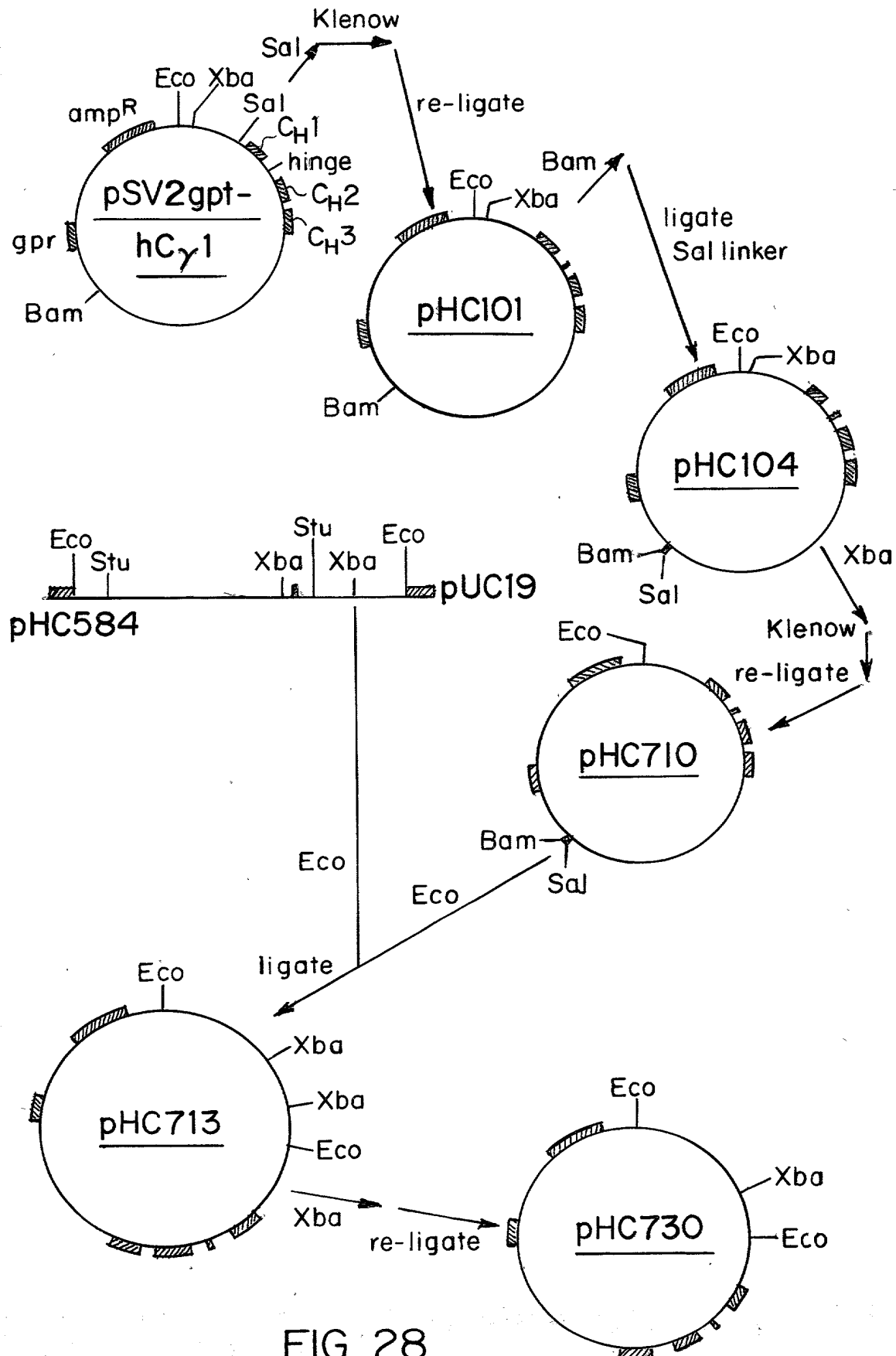


FIG. 28

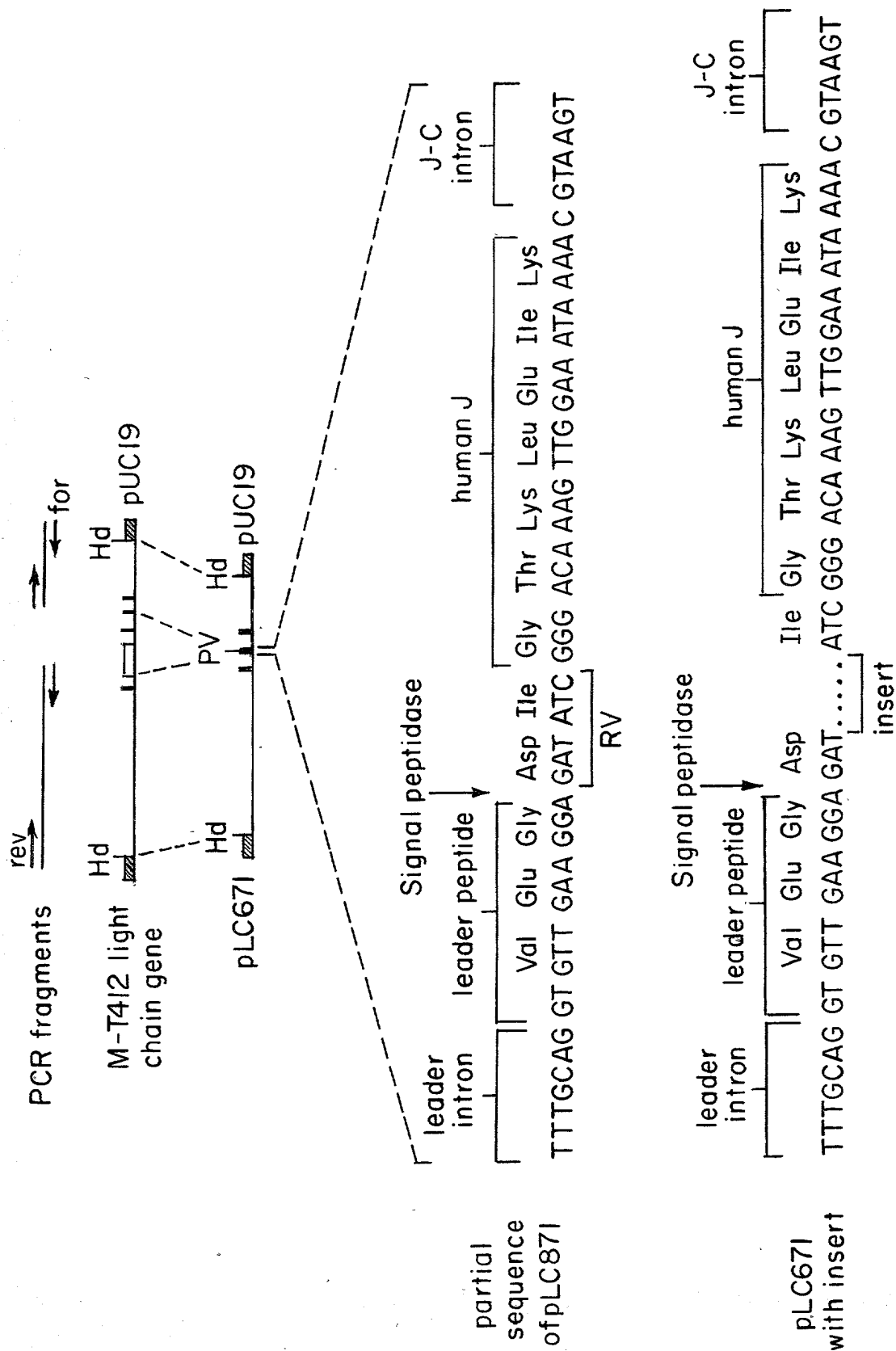


FIG. 29

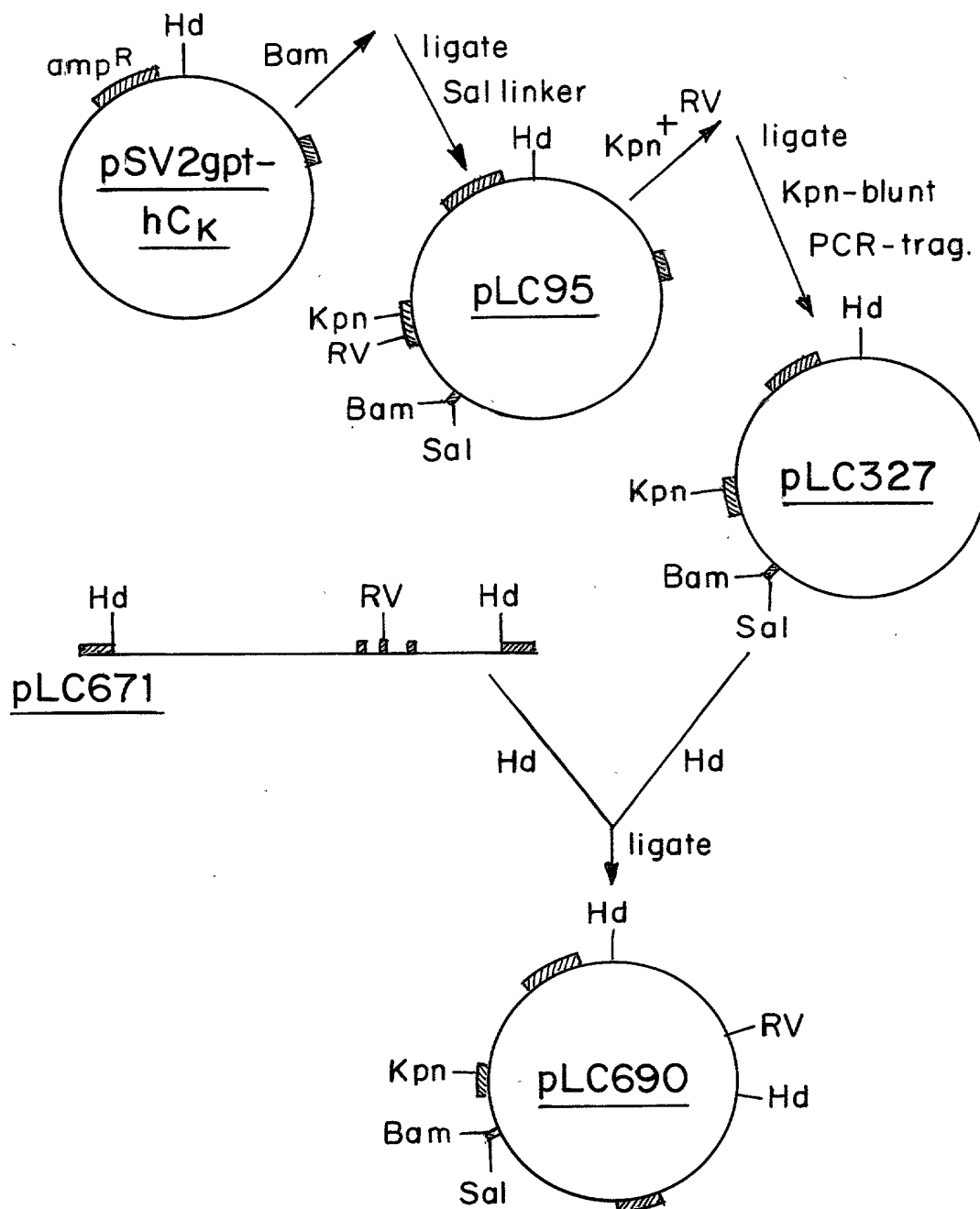


FIG. 30

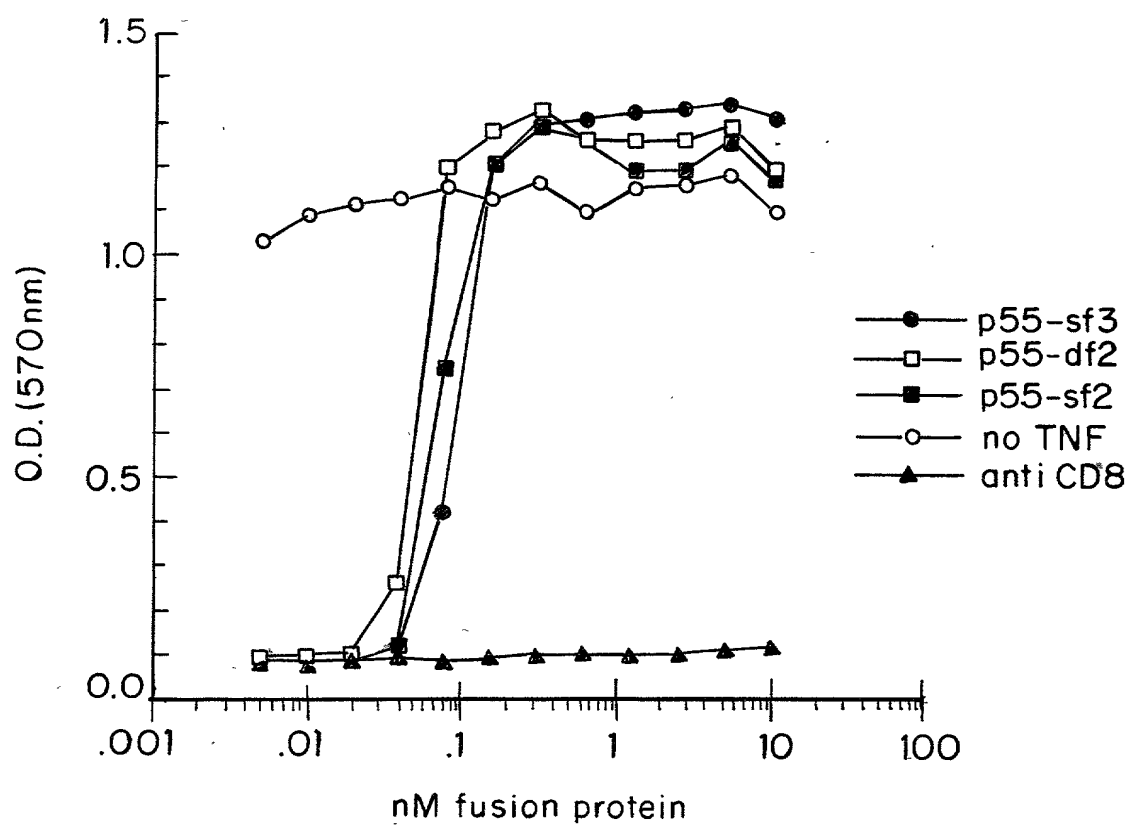


FIG. 31A

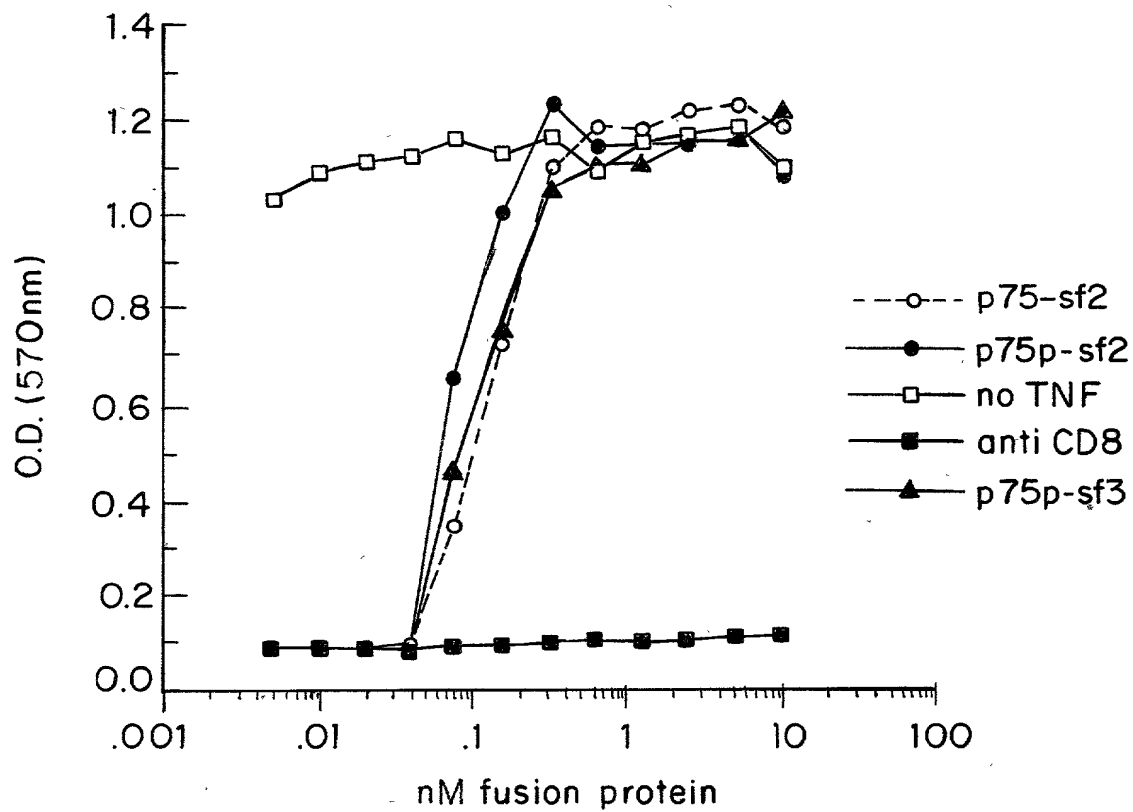


FIG. 3IB

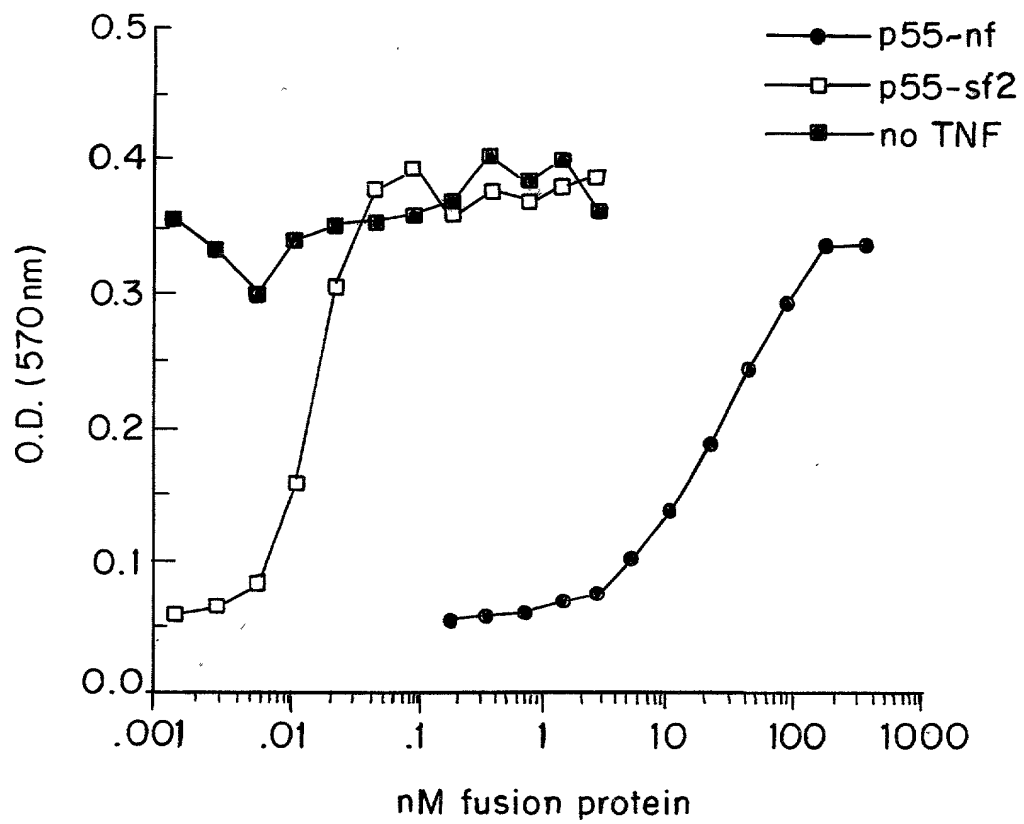


FIG. 31C

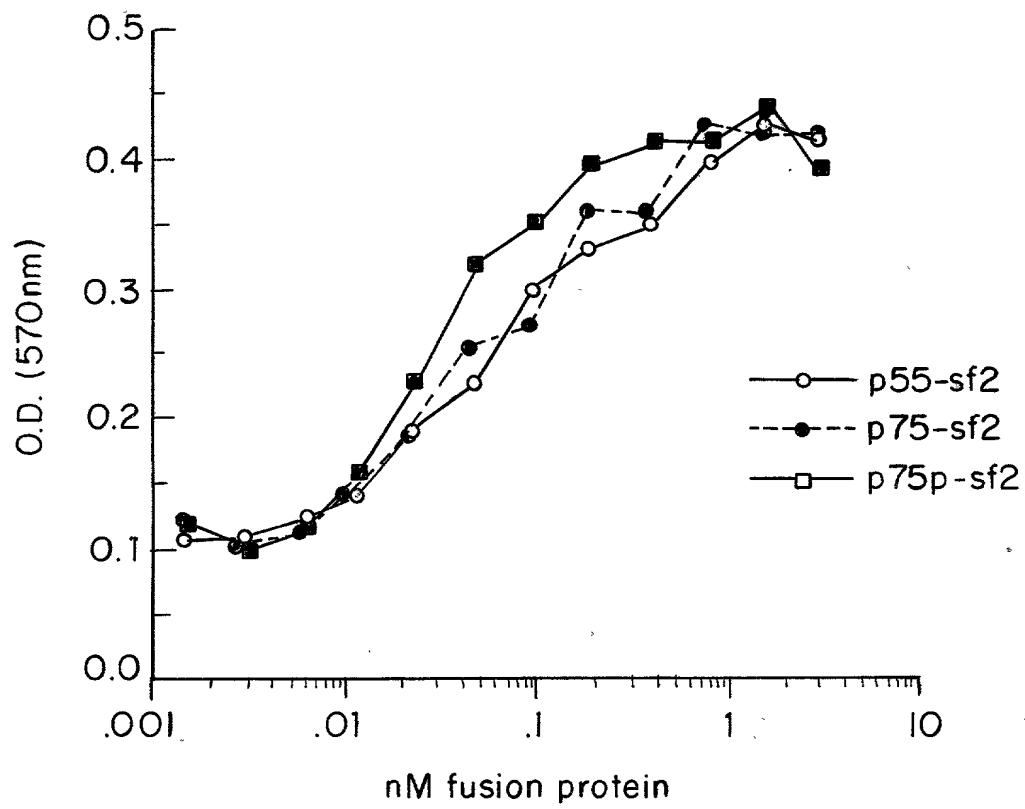


FIG. 32

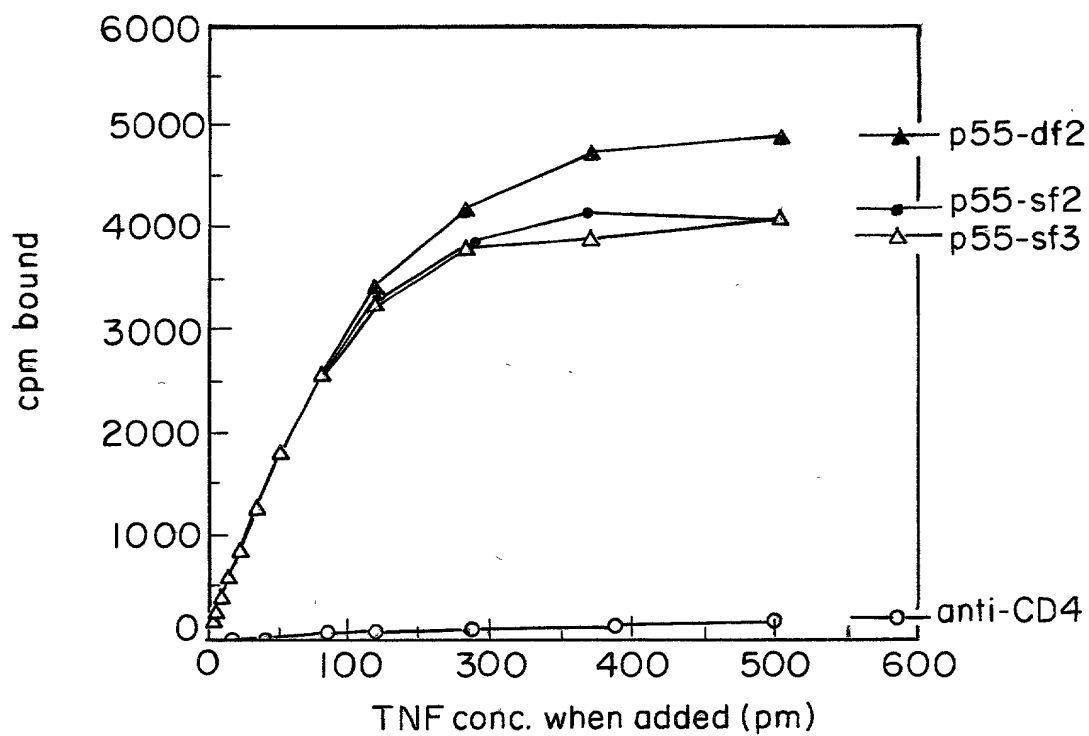


FIG. 33A

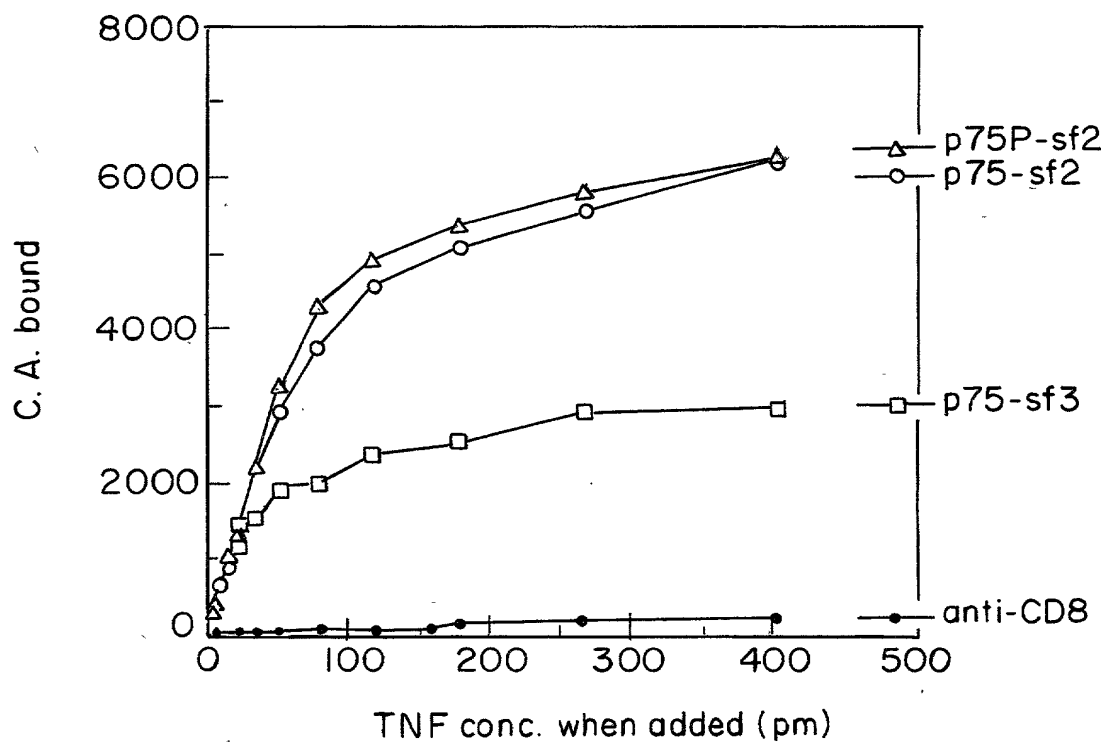


FIG. 33B

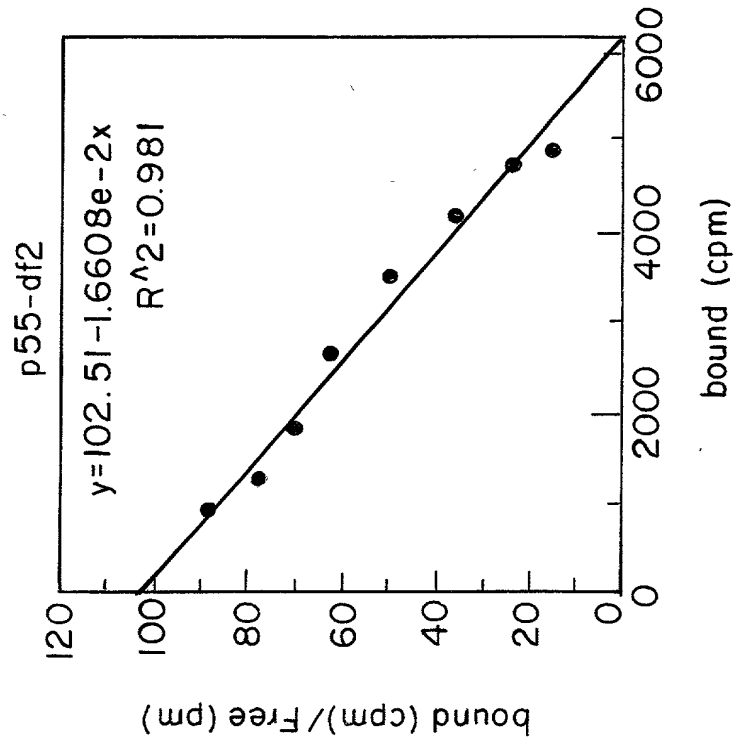


FIG. 33D

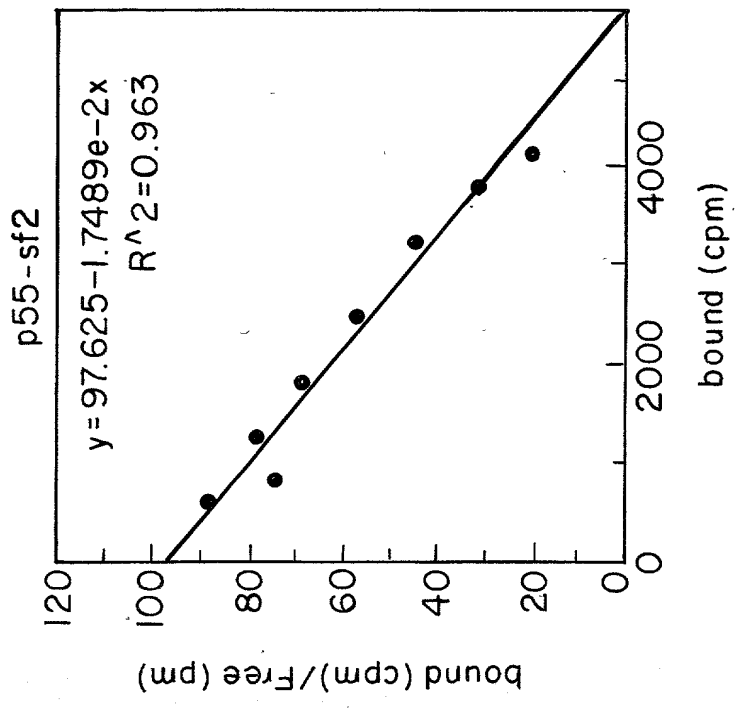


FIG. 33C

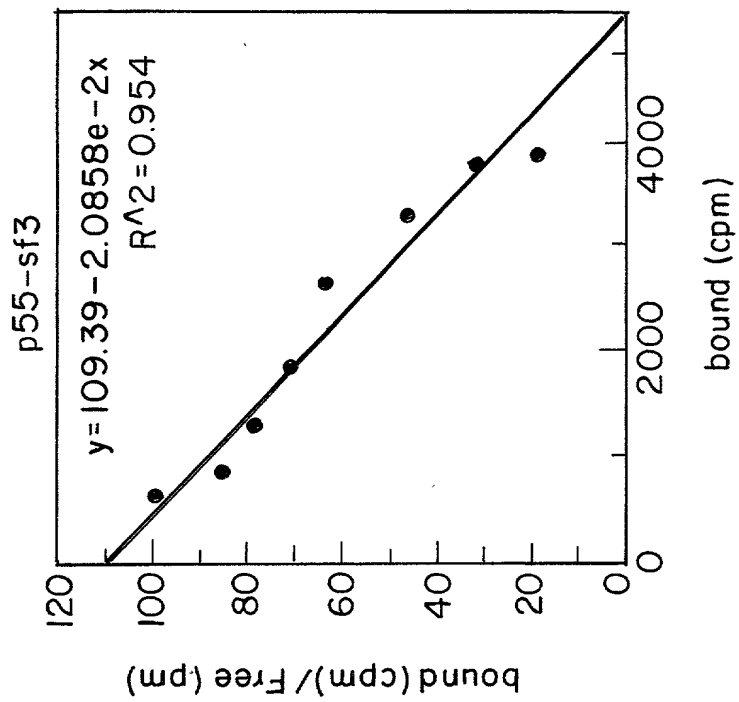


FIG. 33E

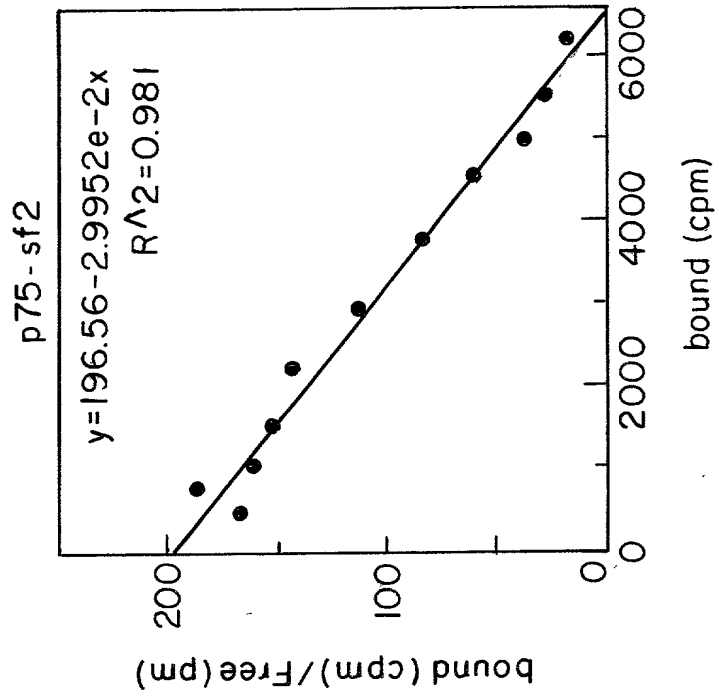


FIG. 33F

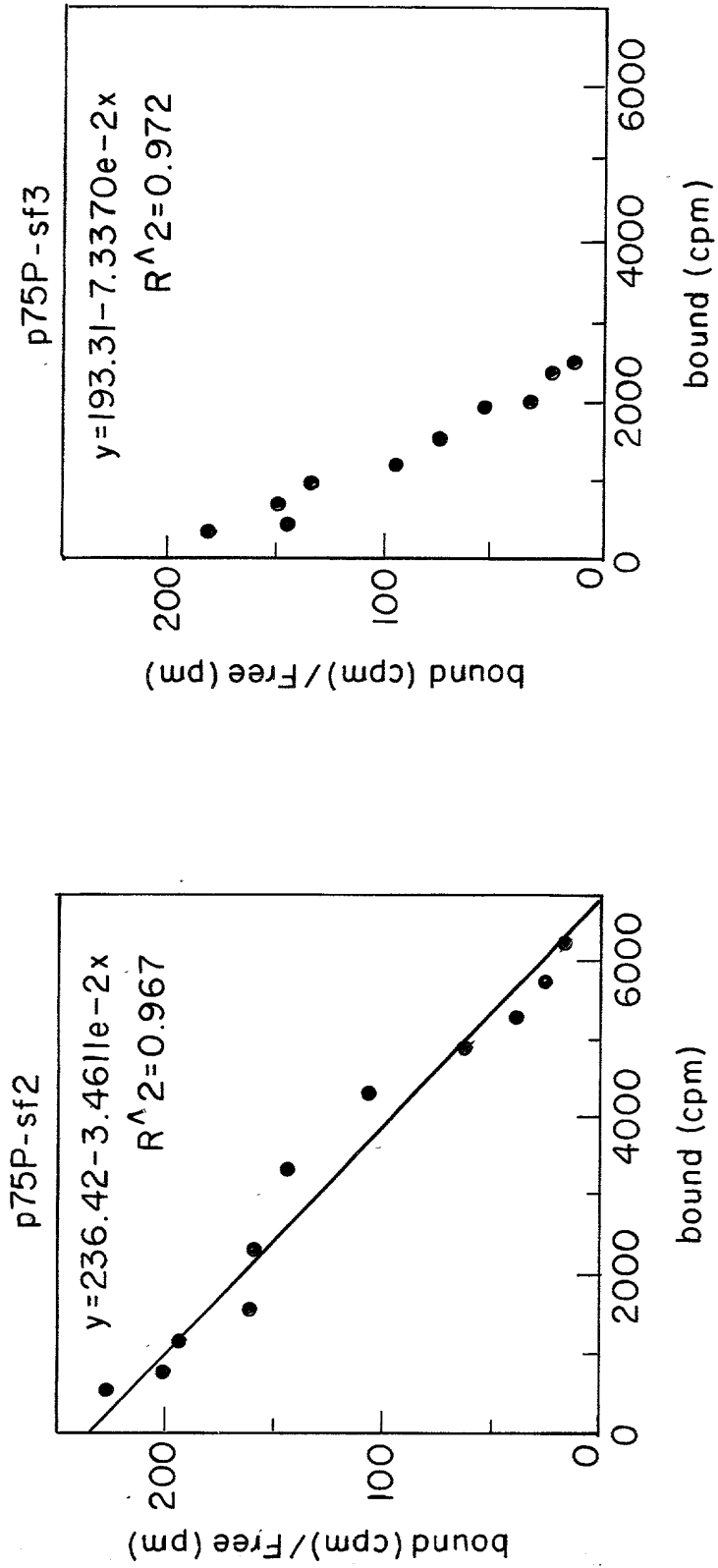
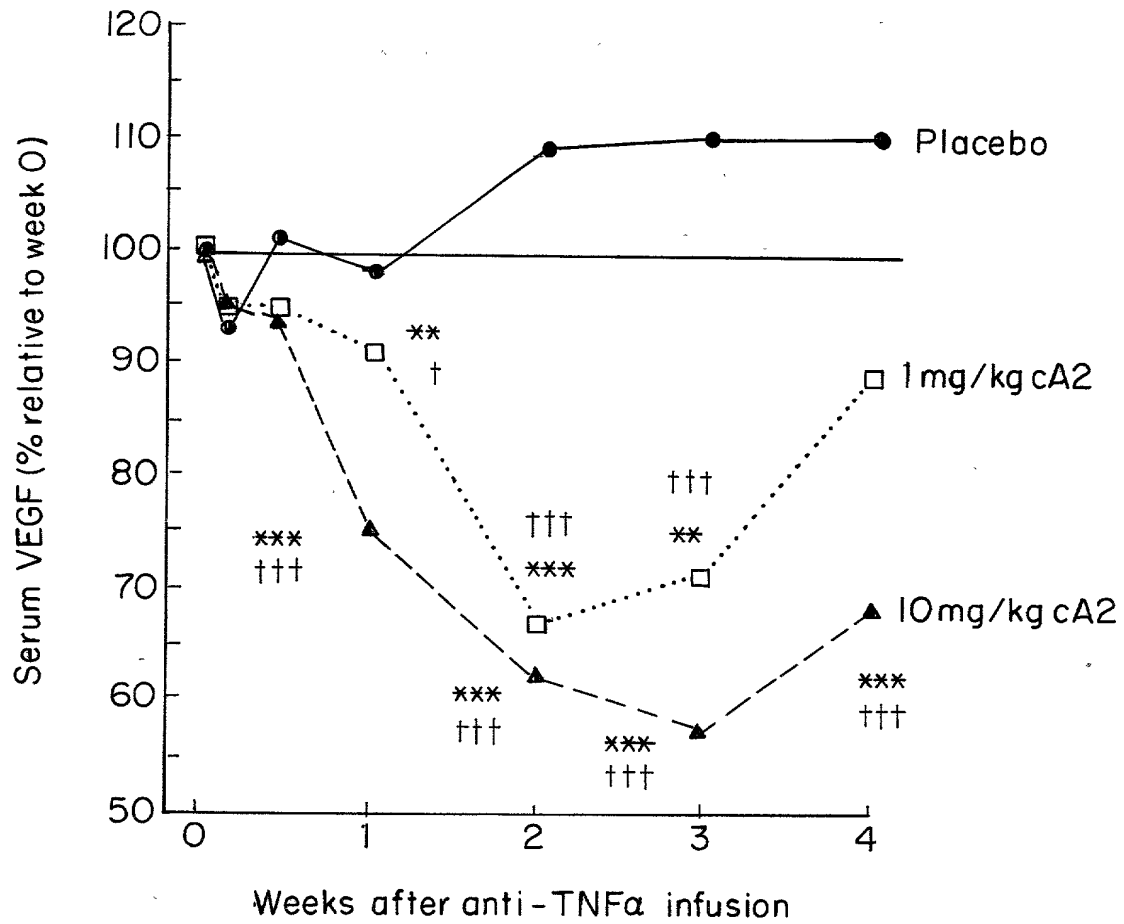


FIG. 33G

FIG. 33H



* p \leq 0.05, ** p \leq 0.01, *** p \leq 0.001 *versus* pre-infusion
 † p \leq 0.05, †† p \leq 0.01, ††† p \leq 0.001 *versus* change in placebo group

FIG. 34